eval simreps

September 2, 2024

```
[3]: import numpy as np
     import os
     from pathlib import Path
     import matplotlib.pyplot as plt
     import tensorflow as tf
     import tensorflow.keras as keras
     from tensorboard.plugins import projector
     import sys
     from PIL import Image
     import smfret.fit as fit
     import smfret.finetuning as finetuning
     from smfret.tf_layers import Attention
     from smfret.tf_layers import Conv
     from smfret.tf_layers import Summary
     from smfret.tf_layers import PrependTaskToken
     from smfret.tf_layers import Embedding
     from smfret.tf_layers import PositionEmbedding
     from smfret.trace simulator import Simulator
     from smfret.trace_simulator import ParameterGenerator
     from smfret.trace_simulator import SimulatedTraceSet
     from smfret.multi_task_learning import FRETStateTraceSet
     from smfret.multi_task_learning import MultistepPhotobleachingTraceSet
     from smfret.multi_task_learning import NoiseLevelTraceSet
     from smfret.dataset import MatlabTraceSet
     from smfret.dataset import FRETTrace
     from smfret.dataset import FRETTraceSet
     from smfret.multi_task_learning import SavedTraceSet
     from sklearn.manifold import TSNE
     from sklearn import metrics
     from sklearn.linear model import LogisticRegression
     from sklearn.linear_model import LinearRegression
```

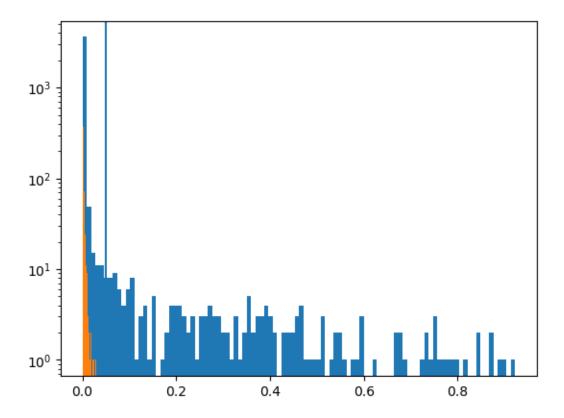
```
from enum import Enum
     import json
     import umap
     import scipy
     from sciplotlib import style as spstyle
     from tqdm.auto import tqdm
[4]: encoder = keras.models.load_model('../saved_models/best_model/
      ⇔encoder-20240511-131505.h5', compile=False)
[5]: def convert_single_channel_data_to_dataset(file, label=None, start_frame=0,_u

max_frames=8000):
         """Converts .MAT data into a model-compatible tensor."""
         key = 'traces'
         conv_width = 100
         dialation factor = 1
         multistep_data = scipy.io.loadmat(file)[key][:, ::dialation_factor]
         n, length = np.shape(multistep_data)
         padding_length = conv_width - length % conv_width
         multistep_data = multistep_data
         padding = np.repeat(multistep_data[:, -2:-1], padding_length, axis=-1)
         multistep_data = np.concatenate([multistep_data, padding], axis=-1)
         time = (1 + np.arange(length + padding_length))
         trace_set = FRETTraceSet()
         trace_set.donor = np.zeros_like(multistep_data)
         trace_set.acceptor = multistep_data
         trace set.time = time
         trace_set.size = n
         if label is None:
             trace_set.label = np.zeros_like(multistep_data)
         else:
             trace_set.label = np.zeros_like(multistep_data) + label
         trace_set.broadcast_data_to_traces()
         # ad-hoc trim
         trace_set.trim(start_frame=start_frame, n_frame=max_frames)
         return trace_set
```

```
[7]: files = '''
../saved_dataset/downstream/simreps/50 fM MUT n1.tif_whole_traces.mat
../saved_dataset/downstream/simreps/50 fM MUT n2.tif_whole_traces.mat
```

```
../saved dataset/downstream/simreps/50 fM MUT n3.tif whole traces.mat
../saved_dataset/downstream/simreps/RE12_NDC-1_whole_traces.mat
../saved_dataset/downstream/simreps/RE12_NDC-2_whole_traces.mat
../saved_dataset/downstream/simreps/RE12_NDC-3_whole_traces.mat
../saved_dataset/downstream/simreps/NDC n1_whole_traces.mat
../saved_dataset/downstream/simreps/NDC n2_whole_traces.mat
../saved_dataset/downstream/simreps/NDC n3_whole_traces.mat
../saved_dataset/downstream/simreps/50 nM WT n1.tif_whole_traces.mat
../saved dataset/downstream/simreps/50 nM WT n2.tif whole traces.mat
../saved_dataset/downstream/simreps/50 nM WT n3.tif_whole_traces.mat
../saved dataset/downstream/simreps/50 nM WT n4.tif whole traces.mat
../saved_dataset/downstream/simreps/50 nM WT n7.tif_whole_traces.mat
'''.strip().split('\n')
trace_sets = []
embedding_list = []
label_list = []
for file in files:
    if 'MUT' in file:
        label = 1
    else:
        label = 0
    trace_set = convert_single_channel_data_to_dataset(file, label,__
 →max frames=1000)
    embedding_list.append(encoder.predict(trace_set.to_tensor()))
    label_list.append(np.max(trace_set.label, axis=-1))
classifier = keras.Sequential([keras.layers.Dense(2)])
classifier.compile(
    optimizer=keras.optimizers.legacy.Adam(learning_rate=1e-2),
    loss=keras.losses.SparseCategoricalCrossentropy(from_logits=True),
)
embedding = np.concatenate(embedding_list, axis=0)
label = np.concatenate(label_list, axis=0)
clf = LogisticRegression(multi_class='multinomial', max_iter=50000, tol=1e-8,__
 →penalty='12', class_weight={0: 2E2, 1: 1})
clf.fit(embedding, label)
6/6 [=======] - 3s 245ms/step
10/10 [======= ] - 2s 191ms/step
7/7 [=======] - 2s 232ms/step
7/7 [=======] - 1s 212ms/step
8/8 [======== ] - 2s 211ms/step
8/8 [=======] - 1s 134ms/step
```

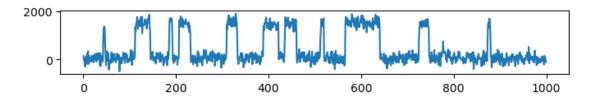
```
8/8 [======== ] - 2s 245ms/step
    6/6 [=======] - 1s 252ms/step
    6/6 [=======] - 1s 261ms/step
    11/11 [======= ] - 1s 134ms/step
    14/14 [======== ] - 2s 179ms/step
    13/13 [========= ] - 2s 191ms/step
    10/10 [======== ] - 1s 135ms/step
    14/14 [========= ] - 3s 185ms/step
[7]: LogisticRegression(class_weight={0: 200.0, 1: 1}, max_iter=50000,
                     multi_class='multinomial', tol=1e-08)
[8]: print(f'Embeddings of {embedding.shape[0]} traces generated.')
    Embeddings of 3904 traces generated.
[12]: | score = clf.predict_proba(embedding)[:, 1]
     WT_cutoff = 0.05
     print('WT Maximum Score', np.max(score[label == 0]))
     print('WT Negative Positive Cutoff', WT_cutoff)
     print('% misclassified WT', np.sum(score[label == 0] >= WT_cutoff) / sum(label_
      \Rightarrow == 0) * 100, '%')
     plt.hist(score, bins=100);
     plt.hist(score[label == 0], bins=100);
     plt.axvline(WT_cutoff)
     plt.yscale('log')
    WT Maximum Score 0.10396381567384576
    WT Negative Positive Cutoff 0.05
    \% misclassified WT 0.031269543464665414 \%
```

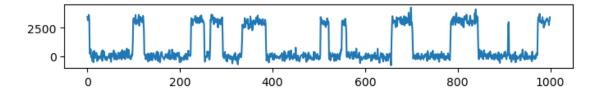


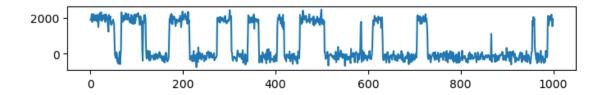
```
[13]: files = '''
      ../saved_dataset/downstream/simreps/50 fM MUT n1.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/50 fM MUT n2.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/50 fM MUT n3.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/1 to 1 mil n1.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/1 to 1 mil n2.tif_whole_traces.mat
      ../saved dataset/downstream/simreps/1 to 1 mil n3.tif whole traces.mat
      ../saved_dataset/downstream/simreps/1 to 100k n1.tif_whole_traces.mat
      ../saved dataset/downstream/simreps/1 to 100k n2.tif whole traces.mat
      ../saved_dataset/downstream/simreps/1 to 100k n3.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/1 to 10k n1.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/1 to 10k n2.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/1 to 10k n3.tif_whole_traces.mat
      ../saved dataset/downstream/simreps/1 to 10k n4.tif whole traces.mat
      ../saved_dataset/downstream/simreps/50 nM WT n5.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/50 nM WT n6.tif_whole_traces.mat
      ../saved_dataset/downstream/simreps/50 nM WT n8.tif_whole_traces.mat
      '''.strip().split('\n')
      test sets = []
      embedding_list = []
      pred_list = []
```

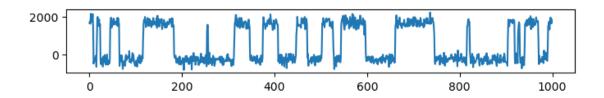
```
for file in files:
    test_set = convert_single_channel_data_to_dataset(file, max_frames=1000)
    embedding_list.append(encoder.predict(test_set.to_tensor()))
    pred_list.append(clf.predict_proba(embedding_list[-1])[:, 1] >= WT_cutoff)
# top score traces
    ranking = np.argsort(clf.predict_proba(embedding_list[-1])[:, 0])
    for i in range(5):
        _, ax = plt.subplots(1, 1, figsize=(8, 1))
        ax.plot(test_set.traces[ranking[i]].acceptor)
    plt.show()
```

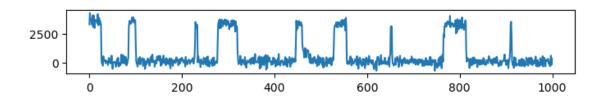
6/6 [======] - 1s 147ms/step



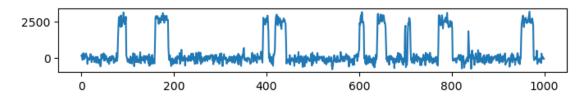


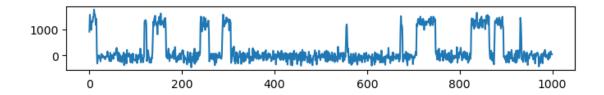


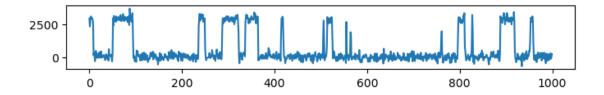


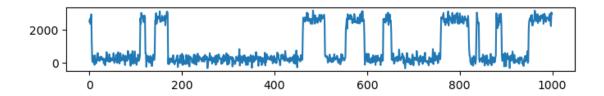


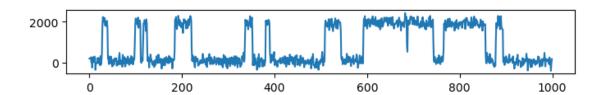
10/10 [======] - 1s 137ms/step



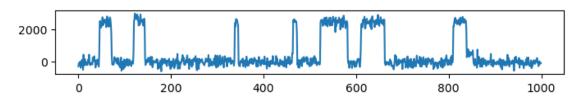


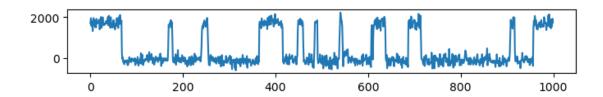


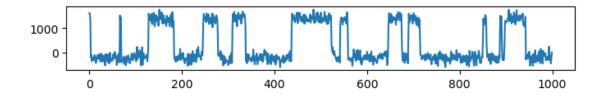


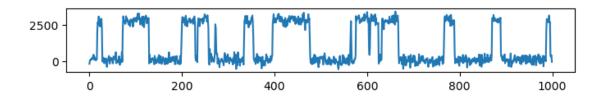


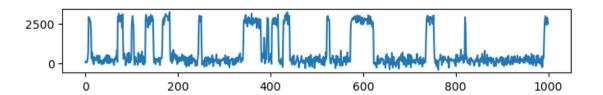
7/7 [======] - 1s 136ms/step



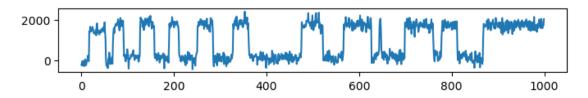


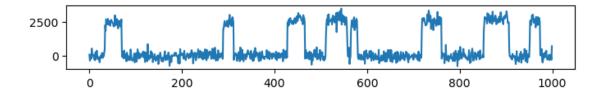


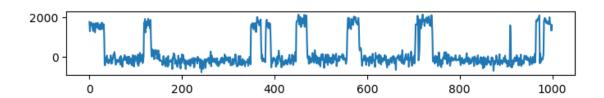


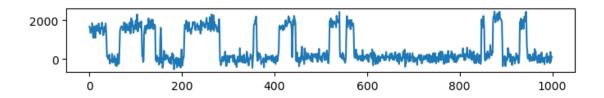


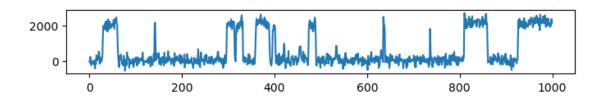
16/16 [=======] - 3s 178ms/step



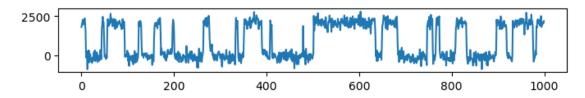


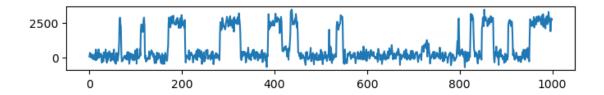


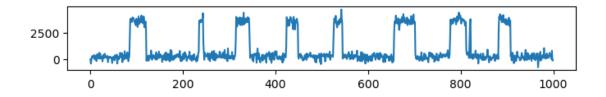


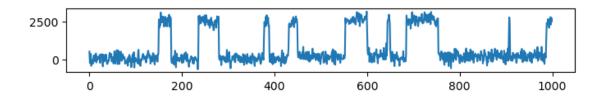


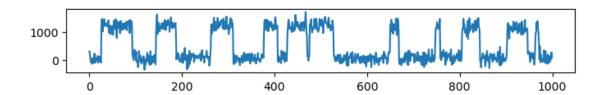
13/13 [=======] - 2s 130ms/step



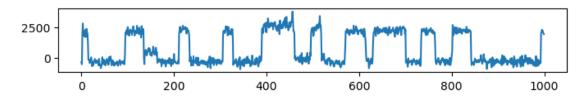


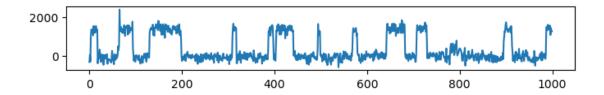


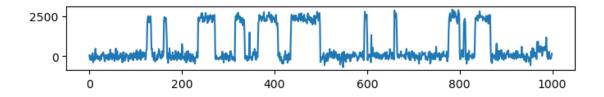


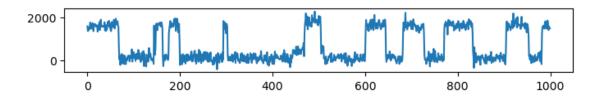


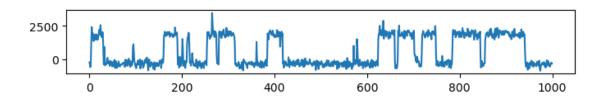
16/16 [=======] - 3s 168ms/step



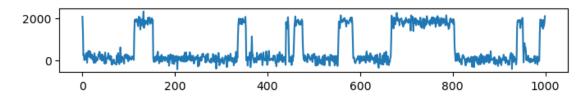


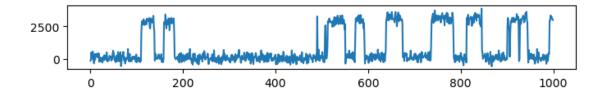


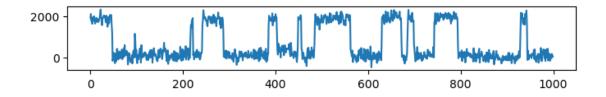


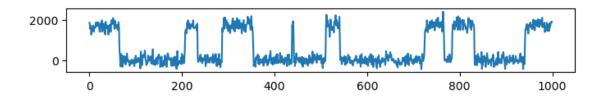


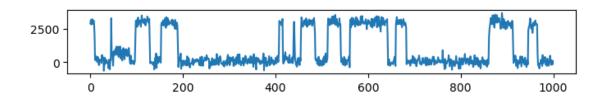
10/10 [======] - 1s 137ms/step



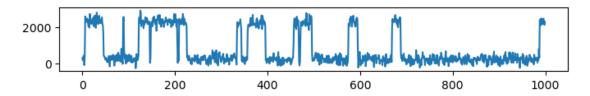


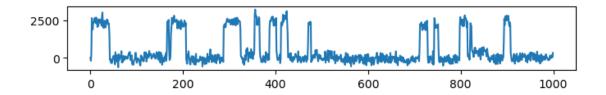


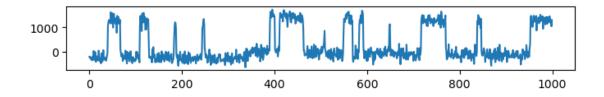


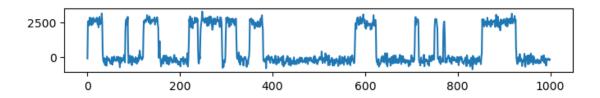


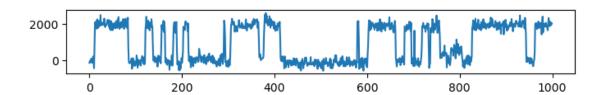
15/15 [=======] - 2s 128ms/step



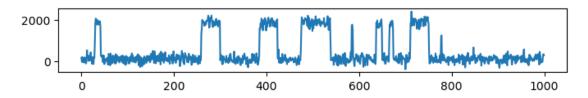


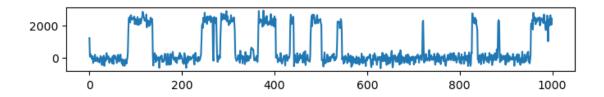


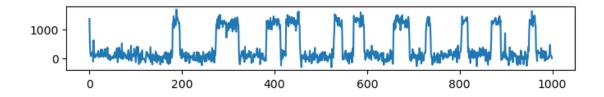


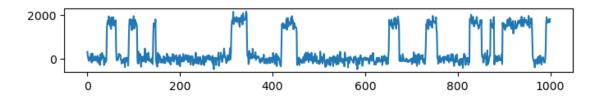


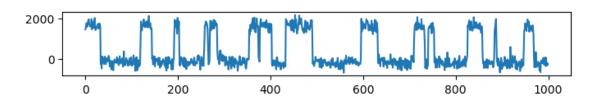
12/12 [======] - 2s 131ms/step



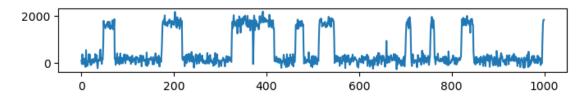


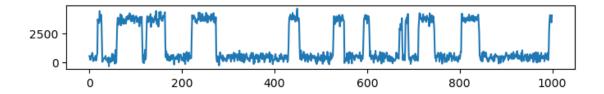


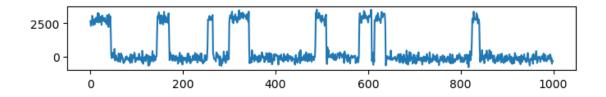


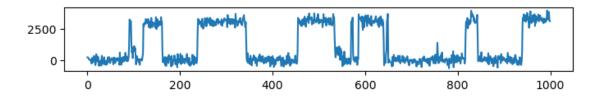


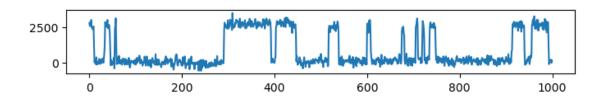
10/10 [======] - 1s 132ms/step



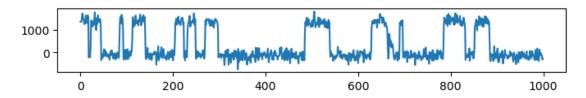


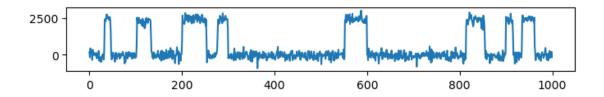


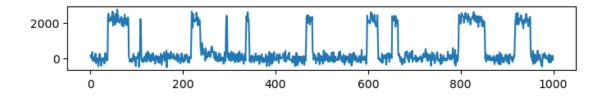


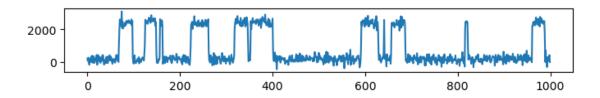


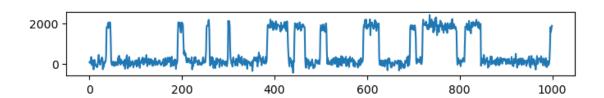
10/10 [======] - 2s 215ms/step



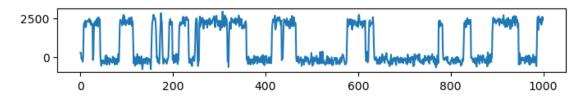


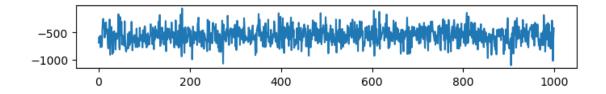


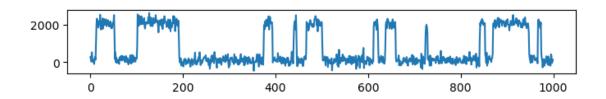


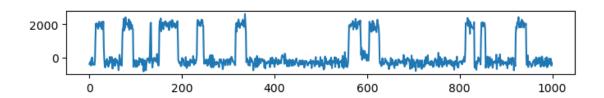


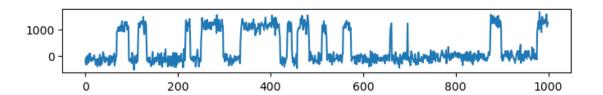
11/11 [======] - 2s 136ms/step



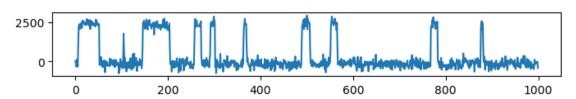


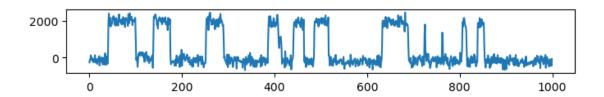


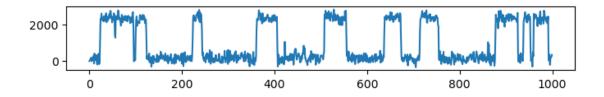


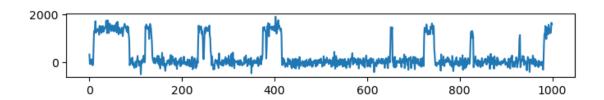


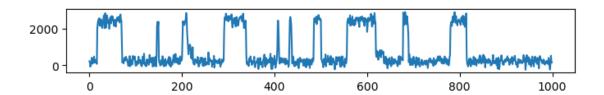




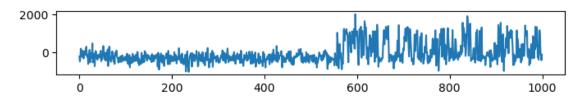


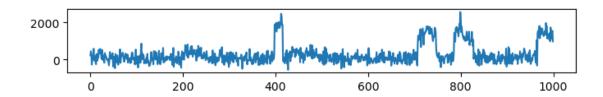


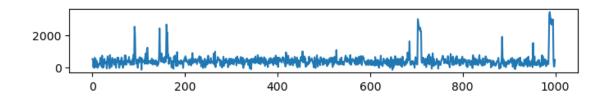


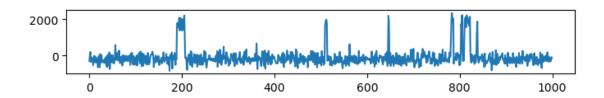


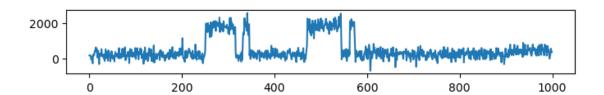
13/13 [======] - 2s 138ms/step

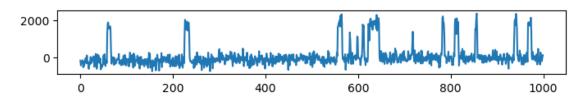


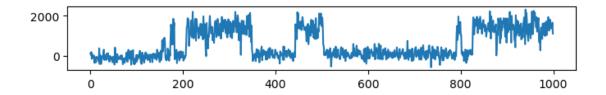


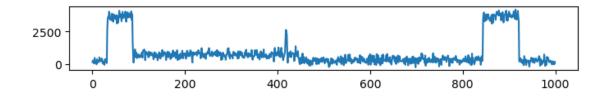


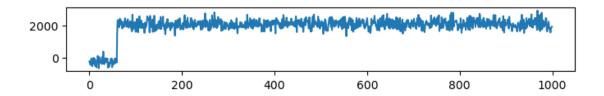


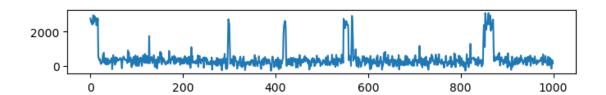




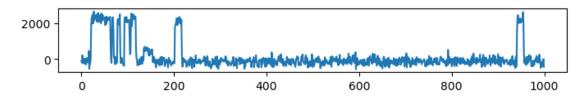


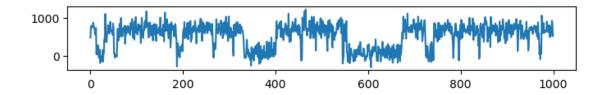


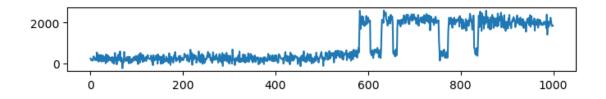


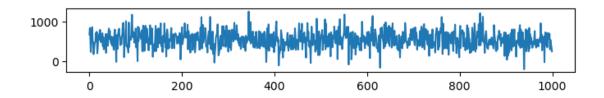


17/17 [========] - 2s 139ms/step









```
5000 - 0 200 400 600 800 1000
```

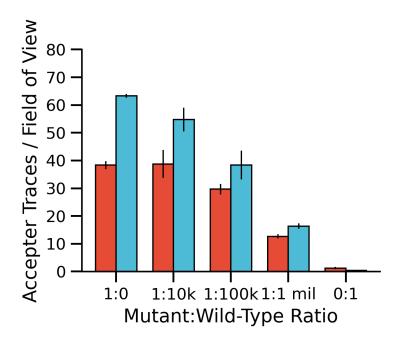
Number of test traces = 5720

```
[18]: groups = ['MUT', '1 to 10k', '1 to 100k', '1 to 1 mil', 'WT']
      names = ['1:0', '1:10k', '1:100k', '1:1 mil', '0:1']
      HMM = [[41, 39, 35], [54, 38, 37, 26], [26, 29, 34], [11, 13, 14], [0, 1, 2, 2]]
      width = 0.36
      with plt.style.context(spstyle.get_style('nature-reviews')):
          color = plt.rcParams['axes.prop_cycle'].by_key()['color']
          fig = plt.figure(figsize=(4, 3))
          plt.xlabel('Mutant:Wild-Type Ratio')
          plt.ylabel('Accepter Traces / Field of View')
          plt.bar(np.arange(len(HMM)) - width / 2, [np.mean(x) for x in HMM],
       ⇔width=width, color=color[0], edgecolor='k', linewidth=1)
          plt.errorbar(np.arange(len(HMM)) - width / 2, [np.mean(x) for x in HMM],
       →[np.std(x) / np.sqrt(len(x)) for x in HMM], linewidth=1, color='black', ___
       ⇔linestyle='none')
          for i, group in enumerate(groups):
              counts = []
              n files = 0
              for j, file in enumerate(files):
                  if group in file:
                      counts.append(np.sum(pred_list[j]))
                      n_files += 1
              count = np.sum(counts) / n_files
              print('ratio between prediction and hmm counting is', count / np.
       →mean(HMM[i]))
              plt.bar([i + width / 2], [count], width=width, color=color[1],__
       ⇔edgecolor='k', linewidth=1)
              plt.errorbar([i + width / 2], [count], [np.std(counts) / np.

sqrt(n_files)], color='black', linewidth=1)
```

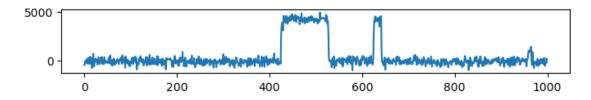
```
plt.xticks(np.arange(len(names)), names)
plt.ylim([0, 80])
plt.savefig('figures/fig3-simreps.svg', transparent=True, dpi=200)
```

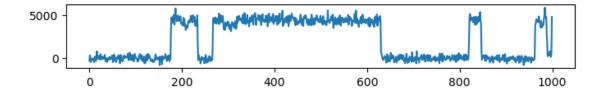
```
ratio between prediction and hmm counting is 1.6521739130434783 ratio between prediction and hmm counting is 1.4129032258064516 ratio between prediction and hmm counting is 1.2921348314606742 ratio between prediction and hmm counting is 1.2894736842105263 ratio between prediction and hmm counting is 0.2666666666666666
```

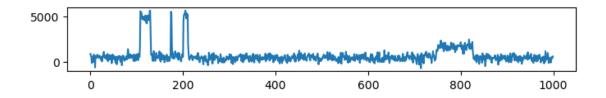


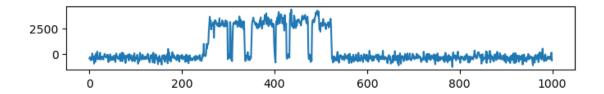
```
[19]: files = '''
      ../saved_dataset/downstream/simreps/RE12_NDC-1_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_NDC-2_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_NDC-3_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_1fM-1_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_1fM-2_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_1fM-3_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_5fM-1_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_5fM-2_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_5fM-3_whole_traces.mat
      ../saved dataset/downstream/simreps/RE12 10fM-2 whole traces.mat
      ../saved_dataset/downstream/simreps/RE12_50fM-1_whole_traces.mat
      ../saved dataset/downstream/simreps/RE12 50fM-2 whole traces.mat
      ../saved_dataset/downstream/simreps/RE12_50fM-3_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_250fM-1_whole_traces.mat
      ../saved_dataset/downstream/simreps/RE12_250fM-2_whole_traces.mat
```

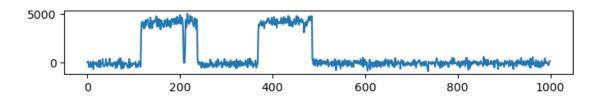
```
../saved_dataset/downstream/simreps/RE12_250fM-3_whole_traces.mat
../saved_dataset/downstream/simreps/RE12_500fM-1_whole_traces.mat
../saved_dataset/downstream/simreps/RE12_500fM-2_whole_traces.mat
../saved_dataset/downstream/simreps/RE12_500fM-3_whole_traces.mat
'''.strip().split('\n')
test_sets = []
embedding_list = []
pred list = []
for file in files:
   print(file)
   test_set = convert_single_channel_data_to_dataset(file, max_frames=1000)
    embedding_list.append(encoder.predict(test_set.to_tensor()))
   pred_list.append(clf.predict_proba(embedding_list[-1])[:, 1] > WT_cutoff)
    # top score traces
   ranking = np.argsort(clf.predict_proba(embedding_list[-1])[:, 0])
   for i in range(5):
        _, ax = plt.subplots(1, 1, figsize=(8, 1))
        ax.plot(test_set.traces[ranking[i]].acceptor)
   plt.show()
```

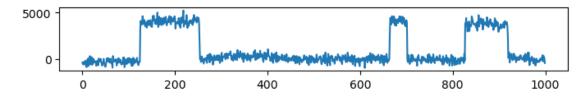



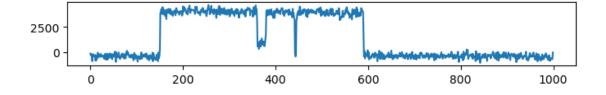


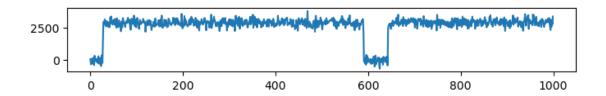


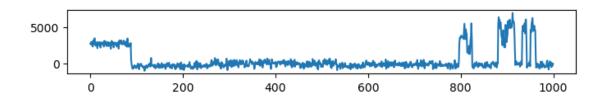


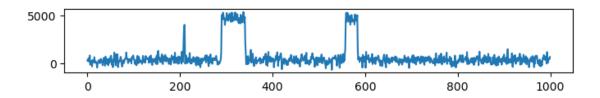


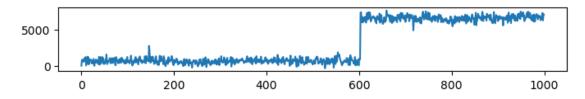


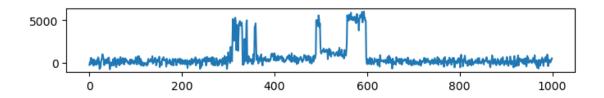


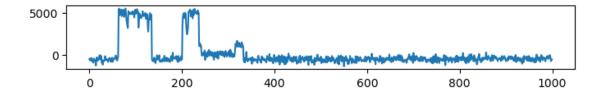


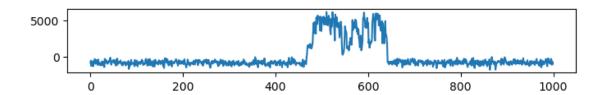


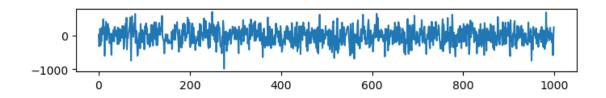


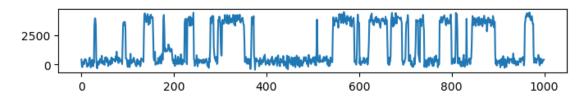


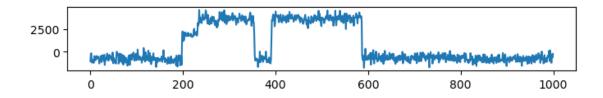


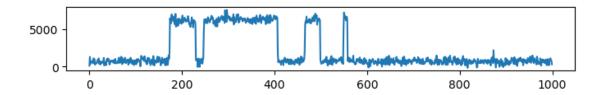


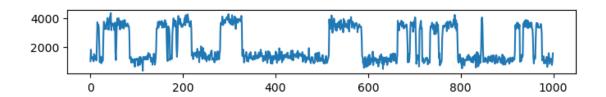


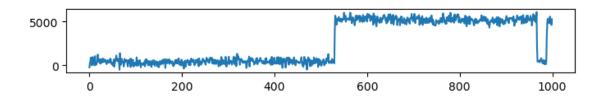


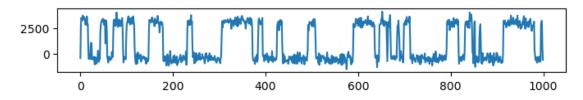


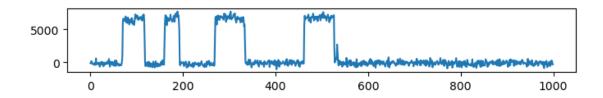


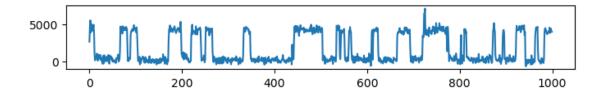


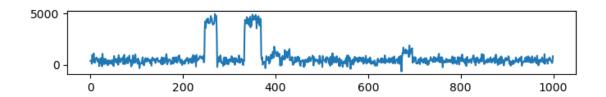


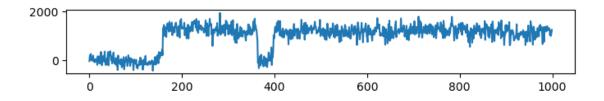


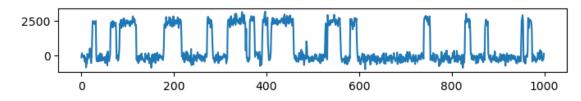


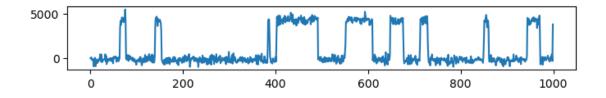


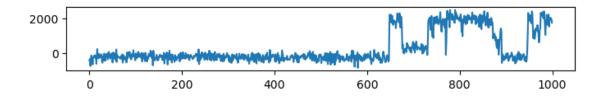


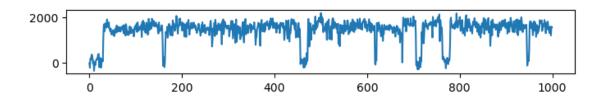


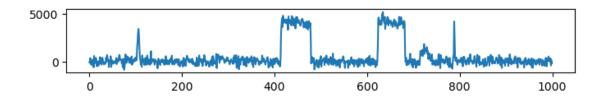


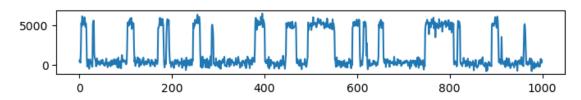


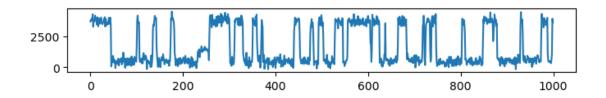


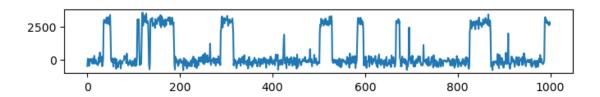


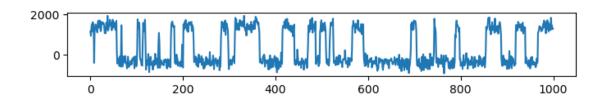


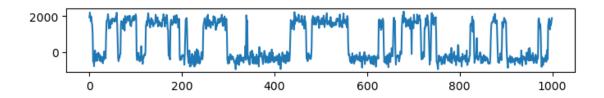


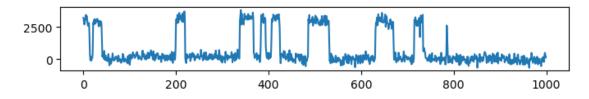


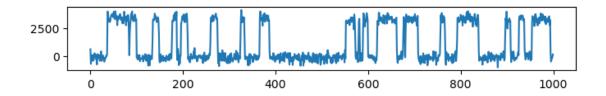


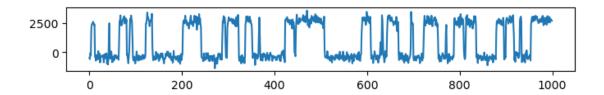


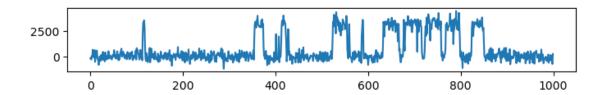


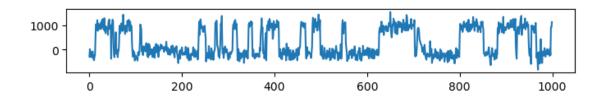


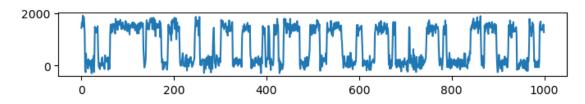


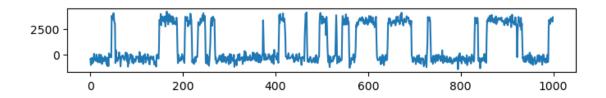


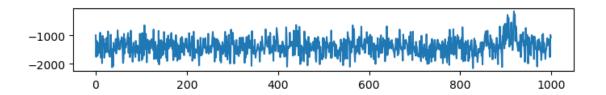


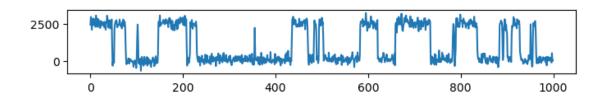


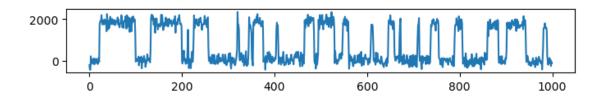




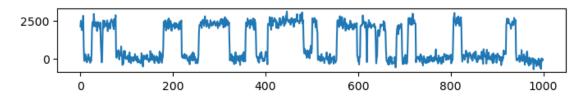


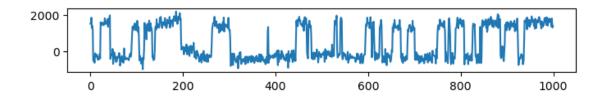


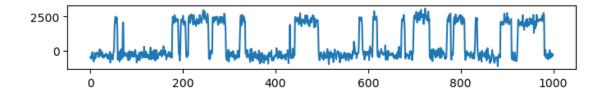


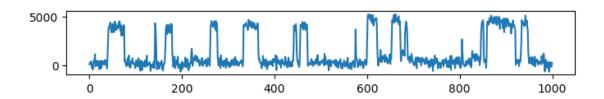


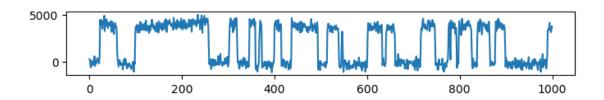
../saved_dataset/downstream/simreps/RE12_10fM-2_whole_traces.mat 17/17 [=========] - 2s 142ms/step

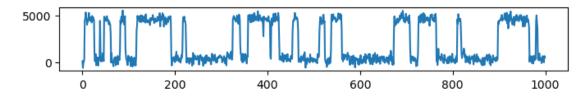


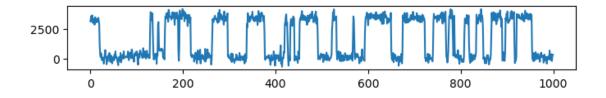


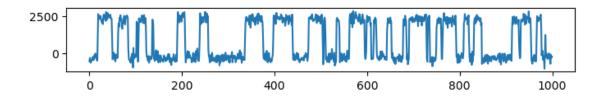


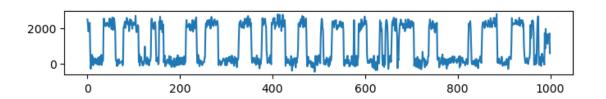


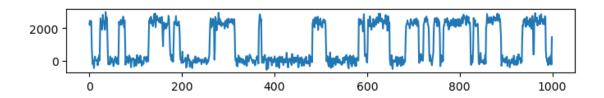




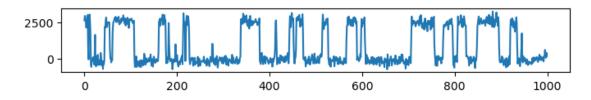


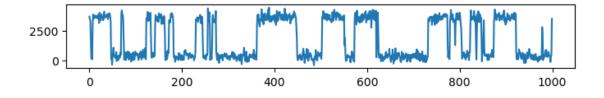


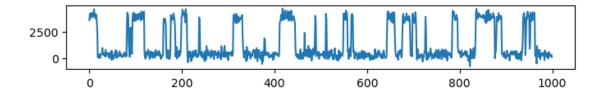


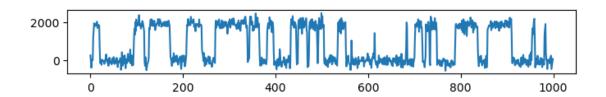


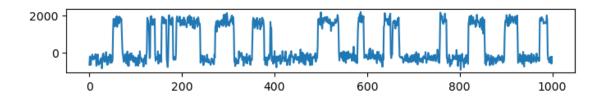
../saved_dataset/downstream/simreps/RE12_50fM-2_whole_traces.mat 9/9 [==========] - 3s 300ms/step



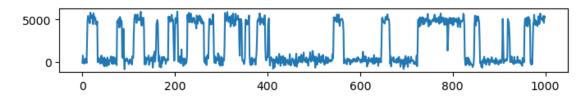


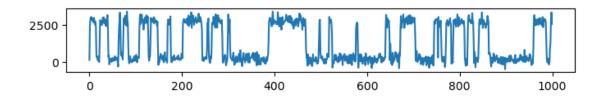


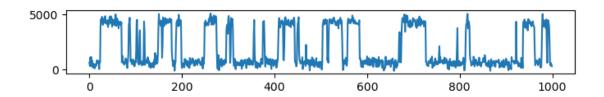


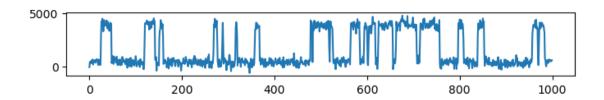


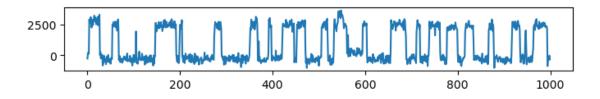
../saved_dataset/downstream/simreps/RE12_50fM-3_whole_traces.mat 5/5 [===========] - 1s 339ms/step

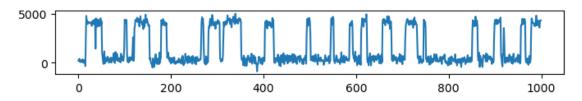


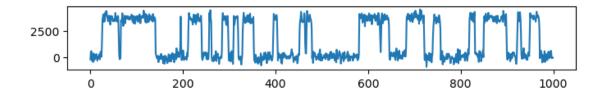


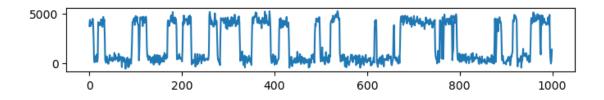


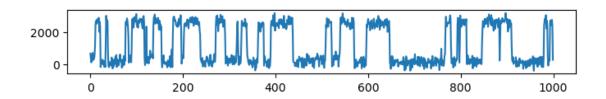


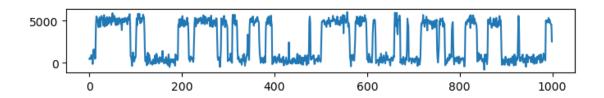


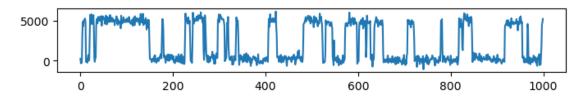


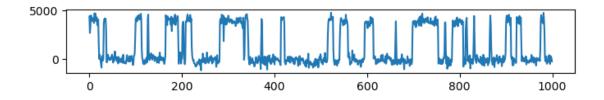


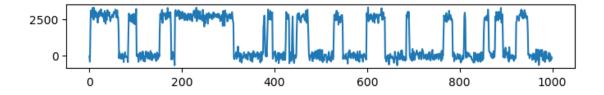


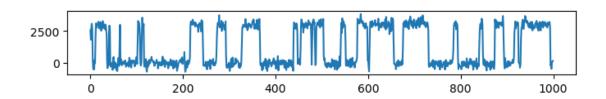


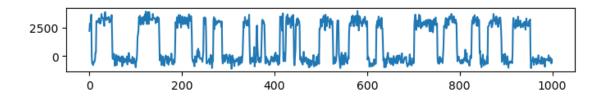


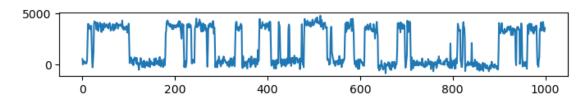


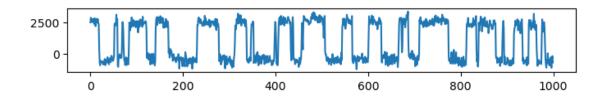


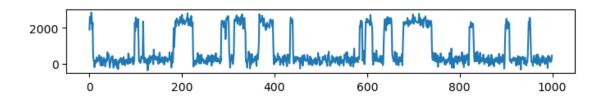


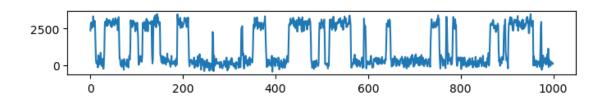


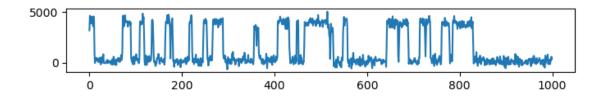


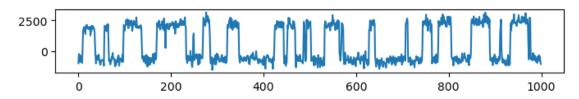


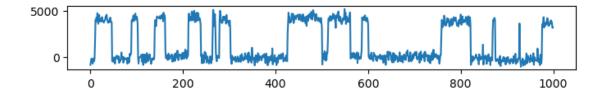


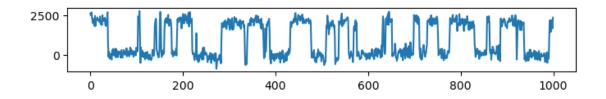


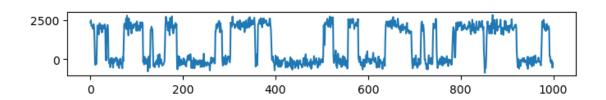


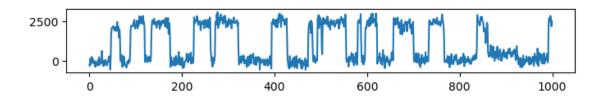


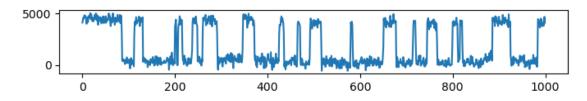


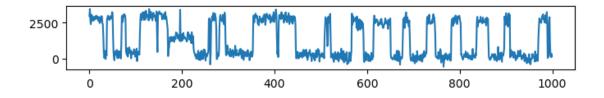


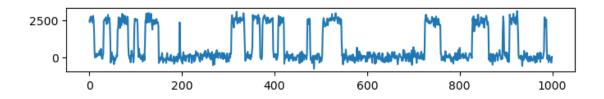


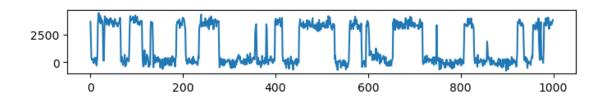


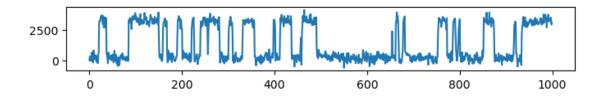


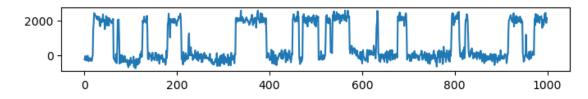


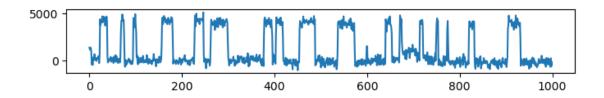


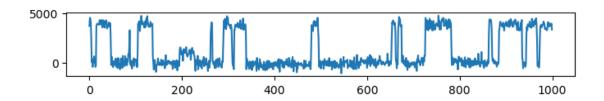


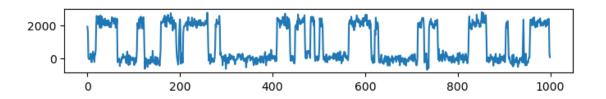


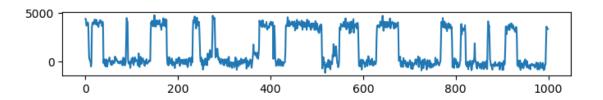












```
[21]: groups = ['_NDC', '_1fM', '_5fM', '_10fM', '_50fM', '_250fM',]
      location = [0, 1, 5, 10, 50, 250,]
      HMM = [0, 0.67, 3.83, 10.33, 57.33, 250.00]
      HMM_err = [0, 0.33, 0.98, 2.33, 2.6, 12.1]
      HMM_reg = LinearRegression(fit_intercept=False).fit(np.expand_dims(location,__
       \Rightarrowaxis=-1), HMM)
      width = 0.36
      with plt.style.context(spstyle.get_style('nature-reviews')):
          color = plt.rcParams['axes.prop_cycle'].by_key()['color']
          fig = plt.figure(figsize=(4, 3))
          plt.xlabel('Concentration (fM)')
          plt.ylabel('Accepter Traces / Field of View')
          plt.scatter(location, HMM, color=color[0], edgecolor=color[0], linewidth=1,__
       ⇔s=20, facecolor='none', marker='d')
          plt.plot(location, HMM_reg.predict(np.expand_dims(location, axis=-1)),__

color=color[0])
          plt.errorbar(location, HMM, HMM_err, color='k', linewidth=1.5,_
       →linestyle='none')
          counts = \Pi
          err = []
          for i, group in enumerate(groups):
              counts.append([])
              n_files = 0
              for j, file in enumerate(files):
                  if group in file:
                      counts[-1].append(np.sum(pred_list[j]))
                      n_files += 1
```

```
count = np.sum(counts[-1]) / n_files
    err.append(np.std(counts[-1]) / np.sqrt(n_files))
    counts[-1] = count

reg = LinearRegression(fit_intercept=False).fit(np.expand_dims(location,u)
axis=-1), counts)

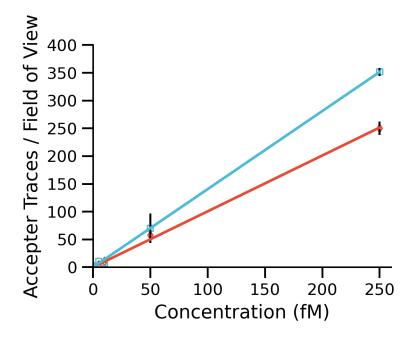
plt.scatter(location, counts, color=color[1], edgecolor=color[1],u)
alinewidth=1, s=20, facecolor='none', marker='s')

plt.plot(location, reg.predict(np.expand_dims(location, axis=-1)),u)
acolor=color[1])

plt.errorbar(location, counts, err, color='k', linewidth=1.5,u)
alinestyle='none')

plt.ylim([0, 400])

plt.xlim([0, 260])
```



```
[27]: print('Increases the sensitivity by', np.round(reg.coef_ / HMM_reg.coef_, Grant of the sensitivity by')
```

Increases the sensitivity by 1.4 fold