Final Project

Course Name: Independent Study - Survival Analysis

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```
In [1]: # Libraries Utilized
        import pandas as pd
        import seaborn as sns
        import matplotlib.pyplot as plt
        from lifelines import KaplanMeierFitter
        from lifelines.statistics import logrank_test, pairwise_logrank_test
        from tabulate import tabulate
        import warnings
        from lifelines import CoxPHFitter
        import matplotlib as mpl
        import numpy as np
        from lifelines import CoxPHFitter
        # !pip install scikit-survival
        from sksurv.ensemble import RandomSurvivalForest
        from sksurv.metrics import concordance_index_censored
        from sklearn.inspection import permutation_importance
        from sksurv.metrics import concordance_index_censored
In [2]: # Loading the CSV file
        canine = pd.read_csv('Canine_Survival_Analysis.csv')
        # Displaying the first few rows
        print(canine.head())
```

```
PatientID site Died how died Pre-ACTH Post-ACTH Insurance weight \
                        0 0.0 0.0 0
       1.0 183.0 0.0
                                                            7.55
       2.0 284.0 0.0
                             0
                                   0.0
                                            0.0
                                                       0
                                                            7.20
1
                             0
2
       3.0 182.0 1.0
                                   267.0
                                            922.0
                                                       0 12.50
3
       4.0 284.0 1.0
                            0
                                  0.0
                                            0.0
                                                       0 19.00
       5.0 159.0 1.0 Euthanasia 145.0
                                            684.0 Insured
                                                           34.25
  BreedRelativeWeight BirthDate ... Complications Hypertensive_Yes4 \
               1.0 14-05-2003 ...
                                      Unknown
0
                                                     Unknown
               1.0 07-11-2002 ...
1
                                      Unknown
                                                     Unknown
2
               2.0 01-02-2005 ...
                                      Unknown
                                                     Unknown
3
               1.0 09-10-2003 ...
                                      Unknown
                                                     Unknown
4
                2.0 01-07-2004 ...
                                         Yes
                                                         Yes
 Oversuppresion? Cortisol stayed <250 Cortisol went <40 \
                             Yes
           Yes
                                             Yes
0
           Yes
1
                         Unknown
                                         Unknown
2
           Yes
                            Yes
                                            No
3
           Yes
                             No
                                             No
4
           Yes
                             Yes
                                             Yes
 Number_comorbidities comorb_UTI comorb_dm comorb_hypot Unnamed: 37
0
               0.0
                         No
                                             No
                                                       NaN
1
                1.0
                         No
                                 No
                                             No
                                                       NaN
2
                3.0
                         No
                                 Yes
                                             No
                                                       NaN
3
                0.0
                        No
                                No
                                             No
                                                       NaN
4
                1.0
                        Yes
                                  No
                                             No
                                                       NaN
```

[5 rows x 38 columns]

Analyzing and Preprocessing - Part 1

```
In [3]: # Printing the column names
        print(canine.columns)
        print(f"Total number of columns: {len(canine.columns)}")
       Index(['PatientID', 'site', 'Died', 'how died', 'Pre-ACTH', 'Post-ACTH',
              'Insurance', 'weight', 'BreedRelativeWeight', 'BirthDate', 'Sex',
              'Isneutered', 'Breed', 'KC_group', 'Purebreed_status',
              'Date of diagnosis_3', 'Failuredate', 'Date of first suspicion_4',
              'Date trilostane started_5', 'Changes to trilostane_6', 'Stay_vs_stop',
              'Treated with trilostane_7', 'Trilostane starting dose (mg/kg)_8',
              'Trilostane SID/BID_9', 'Censored_10', 'Why censored_11',
              'Cause of death_13', 'Neuro signs', 'Complications',
              'Hypertensive_Yes4', 'Oversuppresion?', 'Cortisol stayed <250',
              'Cortisol went <40', 'Number_comorbidities', 'comorb_UTI', 'comorb_dm',
              'comorb hypot', 'Unnamed: 37'],
             dtype='object')
       Total number of columns: 38
In [4]: # Removing the last column
        canine = canine.iloc[:, :-1]
        # Verifying the number of columns after removal
        print(f"Total number of columns after removal: {len(canine.columns)}")
```

```
Total number of columns after removal: 37
In [5]: # Displaying detailed information about the DataFrame
       print(canine.info())
      <class 'pandas.core.frame.DataFrame'>
      RangeIndex: 393 entries, 0 to 392
      Data columns (total 37 columns):
       # Column
                                            Non-Null Count Dtype
      --- -----
                                            _____
       a
          PatientID
                                            219 non-null float64
       1
          site
                                            219 non-null float64
                                            219 non-null float64
          Died
                                           219 non-null object
219 non-null float64
       3
          how died
       4
          Pre-ACTH
       5
          Post-ACTH
                                           219 non-null float64
                                          219 non-null object
       6
          Insurance
                                          219 non-null float64
       7
          weight
          BreedRelativeWeight
                                          219 non-null float64
                                           219 non-null object
       9
           BirthDate
       10 Sex
                                          219 non-null object
       11 Isneutered
                                          219 non-null object
                                          219 non-null object
       12 Breed
                                          219 non-null object
219 non-null float64
       13 KC_group
       14 Purebreed_status
       15 Date of diagnosis_3
                                          219 non-null object
                                          219 non-null object
       16 Failuredate
                                          219 non-null object
       17 Date of first suspicion 4
                                          205 non-null object
       18 Date trilostane started_5
                                          206 non-null float64
       19 Changes to trilostane_6
       20 Stay_vs_stop
                                          219 non-null float64
                                            219 non-null float64
       21 Treated with trilostane_7
       22 Trilostane starting dose (mg/kg)_8 199 non-null float64
       23 Trilostane SID/BID_9
                                            206 non-null object
                                           45 non-null object
       24 Censored_10
       25 Why censored_11
                                           40 non-null
                                                         object
                                           177 non-null object
       26 Cause of death 13
                                          219 non-null object
       27 Neuro signs
                                          219 non-null object
       28 Complications
       29 Hypertensive Yes4
                                          219 non-null object
       30 Oversuppresion?
                                          219 non-null object
       31 Cortisol stayed <250
                                          219 non-null object
       32 Cortisol went <40
                                          219 non-null object
                                          219 non-null float64
       33 Number comorbidities
       34 comorb_UTI
                                          219 non-null object
       35 comorb_dm
                                           219 non-null object
       36 comorb_hypot
                                            219 non-null object
      dtypes: float64(13), object(24)
      memory usage: 113.7+ KB
      None
In [6]: # Creating a subset excluding PatientID
       canine_subset = canine.iloc[:219].drop('PatientID', axis=1)
```

```
canine_subset = canine.iloc[:219].drop('PatientID', axis=1)

# Total columns in subset
print(f"Total number of columns in the subset: {len(canine_subset.columns)}")
```

```
# Total rows in subset
print(f"Number of rows in canine_subset: {len(canine_subset)}")
print(canine_subset.head())
print(canine_subset.tail())

# Converting the 'Died' column from float to int
canine_subset['Died'] = canine_subset['Died'].astype(int)

# Printing the unique and the first few values to confirm the conversion
print("Unique values in 'Died' column:", canine_subset['Died'].unique())
print(canine_subset['Died'].head())
```

```
Total number of columns in the subset: 36
Number of rows in canine subset: 219
   site Died
               how died Pre-ACTH Post-ACTH Insurance weight \
          0.0
0 183.0
                               0.0
                                                       7.55
                       0
                                         0.0
                                                        7.20
1 284.0
        0.0
                       0
                               0.0
                                         0.0
                                                     0
2 182.0 1.0
                       0
                             267.0
                                       922.0
                                                     0
                                                       12.50
3 284.0
        1.0
                       0
                               0.0
                                         0.0
                                                       19.00
4 159.0 1.0 Euthanasia
                             145.0
                                                        34.25
                                       684.0
                                               Insured
  BreedRelativeWeight BirthDate
                                   Sex ... Neuro signs Complications
                  1.0 14-05-2003
                                   male ...
                                                     No
                                                              Unknown
0
                 1.0 07-11-2002 Female ...
                                                              Unknown
1
                                                     No
                  2.0 01-02-2005
2
                                   male
                                                     No
                                                              Unknown
                                         . . .
3
                  1.0 09-10-2003 Female ...
                                                              Unknown
4
                  2.0 01-07-2004
                                   male ...
                                                      No
                                                                 Yes
 Hypertensive_Yes4 Oversuppresion? Cortisol stayed <250 Cortisol went <40 \
           Unknown
                               Yes
0
                                                   Yes
                                                                    Yes
           Unknown
                               Yes
                                               Unknown
                                                                Unknown
2
           Unknown
                               Yes
                                                   Yes
                                                                    No
           Unknown
3
                               Yes
                                                                    No
                                                    No
4
              Yes
                               Yes
                                                   Yes
                                                                    Yes
 Number_comorbidities comorb_UTI comorb_dm comorb_hypot
                  0.0
                             No
1
                  1.0
                             No
                                       No
                                                     No
2
                  3.0
                             No
                                       Yes
                                                     No
3
                  0.0
                            No
                                       No
                                                     No
4
                  1.0
                            Yes
                                       No
                                                     No
[5 rows x 36 columns]
     site Died
                  how died Pre-ACTH Post-ACTH Insurance weight \
214 139.0 1.0 Euthanasia
                            0.0
                                           0.0 Uninsured
                                                            9.10
215 138.0 1.0 Euthanasia
                                0.0
                                           0.0 Uninsured
                                                          9.00
                            118.0
216 120.0 1.0 Unassisted
                                        1380.0 Uninsured
                                                          9.20
217 166.0 0.0 0
                             0.0
                                           0.0 Uninsured 7.95
                            178.0
218 123.0 1.0 Euthanasia
                                         811.0 Insured
                                                          0.00
    BreedRelativeWeight BirthDate
                                     Sex ... Neuro signs Complications ∖
214
                   1.0 26-07-2003
                                     male ...
                                                       No
                                                                Unknown
                   1.0 08-03-2001
                                   male ...
215
                                                      Yes
                                                                Unknown
216
                   1.0 09-06-1995 Female ...
                                                      No
                                                                Unknown
                   4.0 29-09-2004 Female ...
217
                                                      No
                                                                Unknown
                   4.0 03-11-1999
                                                                Unknown
218
                                   male ...
                                                      Yes
   Hypertensive_Yes4 Oversuppresion? Cortisol stayed <250 Cortisol went <40 \
214
             Unknown
                                  No
                                                      No
                                                                       No
215
             Unknown
                                  No
                                                 Unknown
                                                                  Unknown
216
             Unknown
                                  No
                                                    Yes
                                                                       No
217
             Unknown
                                  No
                                                 Unknown
                                                                  Unknown
                                  No
218
             Unknown
                                                 Unknown
                                                                  Unknown
   Number_comorbidities comorb_UTI comorb_dm comorb_hypot
214
                   0.0
                                                      No
                               No
                                         No
215
                   1.0
                               No
                                         No
216
                   0.0
                               No
                                                      No
                                         No
```

```
217
                            0.0
                                        No
                                                   No
                                                                 No
       218
                            1.0
                                        No
                                                   No
                                                                 No
       [5 rows x 36 columns]
       Unique values in 'Died' column: [0 1]
       1
            0
       2
            1
       3
            1
       4
            1
       Name: Died, dtype: int32
In [7]: # Calculating the frequency table for 'site' column
        site_freq = canine_subset['site'].value_counts()
        site_percent = canine_subset['site'].value_counts(normalize=True, dropna=False) * 1
        print("Frequency table for 'site':")
        print("Value Frequency Percentage")
        for value, count in site_freq.items():
            percentage = site_percent[value]
            print(f"{value:<6} {count:<10} {percentage:.2f}%")</pre>
        print(f"\nTotal number of entries: {len(canine_subset)}")
        print(f"Number of unique values: {canine_subset['site'].nunique()}")
        print(f"Number of missing values: {canine_subset['site'].isnull().sum()}")
```

| Frequency table for 'site': | | | | |
|-----------------------------|---|------------|--|--|
| | - | Percentage | | |
| 174.0 | 7 | 3.20% | | |
| 120.0 | | 3.20% | | |
| 118.0 | | 3.20% | | |
| 132.0 | | 2.74% | | |
| | | | | |
| 160.0 | | 2.74% | | |
| 196.0 | | 2.74% | | |
| | 6 | 2.74% | | |
| | 6 | 2.74% | | |
| | 5 | 2.28% | | |
| 178.0 | | 2.28% | | |
| 123.0 | | 2.28% | | |
| 115.0 | | 2.28% | | |
| 140.0 | | 2.28% | | |
| 173.0 | | 1.83% | | |
| 199.0 | 4 | 1.83% | | |
| 133.0 | 4 | 1.83% | | |
| 131.0 | 4 | 1.83% | | |
| 136.0 | 4 | 1.83% | | |
| 163.0 | 4 | 1.83% | | |
| 197.0 | 4 | 1.83% | | |
| 189.0 | 4 | 1.83% | | |
| 151.0 | 4 | 1.83% | | |
| 164.0 | 4 | 1.83% | | |
| 157.0 | 3 | 1.37% | | |
| 107.0 | 3 | 1.37% | | |
| 122.0 | | 1.37% | | |
| 111.0 | | 1.37% | | |
| 145.0 | | 1.37% | | |
| 138.0 | | 1.37% | | |
| 108.0 | | 1.37% | | |
| 121.0 | 3 | 1.37% | | |
| 139.0 | | 1.37% | | |
| 169.0 | 3 | 1.37% | | |
| 144.0 | 3 | 1.37% | | |
| 106.0 | | 1.37% | | |
| 166.0 | | 1.37% | | |
| 159.0 | 3 | 1.37% | | |
| 110.0 | 3 | 1.37% | | |
| | 2 | | | |
| 183.0 | | 0.91% | | |
| 168.0 | 2 | 0.91% | | |
| 158.0 | 2 | 0.91% | | |
| 187.0 | 2 | 0.91% | | |
| 176.0 | | 0.91% | | |
| 113.0 | | 0.91% | | |
| 186.0 | 2 | 0.91% | | |
| 143.0 | 2 | 0.91% | | |
| 162.0 | 2 | 0.91% | | |
| 154.0 | 2 | 0.91% | | |
| 128.0 | 2 | 0.91% | | |
| 165.0 | 2 | 0.91% | | |
| 182.0 | 2 | 0.91% | | |
| 124.0 | 2 | 0.91% | | |
| 284.0 | 2 | 0.91% | | |
| 134.0 | 2 | 0.91% | | |
| | | | | |

```
194.0 2
                        0.91%
       152.0 2
                        0.91%
       293.0 1
                        0.46%
       127.0 1
                        0.46%
       147.0 1
                        0.46%
       172.0 1
                        0.46%
      137.0 1
                        0.46%
       142.0 1
                        0.46%
       117.0 1
                        0.46%
       179.0 1
                        0.46%
       170.0 1
                        0.46%
       292.0 1
                        0.46%
       155.0 1
                        0.46%
       295.0 1
                        0.46%
       148.0 1
                        0.46%
      156.0 1
                        0.46%
      167.0 1
                        0.46%
       294.0 1
                      0.46%
       296.0 1
                        0.46%
       188.0 1
                        0.46%
       146.0 1
                        0.46%
       181.0 1
                      0.46%
       114.0 1
                      0.46%
       130.0 1
                      0.46%
       Total number of entries: 219
       Number of unique values: 78
       Number of missing values: 0
In [8]: # "site" column not so informative, thus dropping from the analysis
        canine_subset = canine_subset.iloc[:, 1:]
        # Total columns in subset
        print(f"Total number of columns in the subset: {len(canine_subset.columns)}")
        # Colmun names in subset
        print(f"Column names in the subset: {canine_subset.columns}")
       Total number of columns in the subset: 35
       Column names in the subset: Index(['Died', 'how died', 'Pre-ACTH', 'Post-ACTH', 'Ins
       urance', 'weight',
              'BreedRelativeWeight', 'BirthDate', 'Sex', 'Isneutered', 'Breed',
              'KC_group', 'Purebreed_status', 'Date of diagnosis_3', 'Failuredate',
              'Date of first suspicion_4', 'Date trilostane started_5',
              'Changes to trilostane_6', 'Stay_vs_stop', 'Treated with trilostane_7',
              'Trilostane starting dose (mg/kg)_8', 'Trilostane SID/BID_9',
              'Censored_10', 'Why censored_11', 'Cause of death_13', 'Neuro signs',
              'Complications', 'Hypertensive_Yes4', 'Oversuppresion?',
              'Cortisol stayed <250', 'Cortisol went <40', 'Number_comorbidities',
              'comorb_UTI', 'comorb_dm', 'comorb_hypot'],
             dtype='object')
In [9]: # Printing the count of NaN values in each column
        print("Count of NaN values in each column before replacement:")
        print(canine subset.isnull().sum())
        # There were a total of 393 rows initially.
        # Removing the nulls resulted in a dataset of 219 rows
```

```
Count of NaN values in each column before replacement:
        Died
        how died
                                                 0
        Pre-ACTH
                                                 0
        Post-ACTH
                                                 0
        Insurance
                                                 0
        weight
                                                 0
        BreedRelativeWeight
                                                 0
                                                 0
        BirthDate
        Sex
                                                 0
        Isneutered
                                                 0
        Breed
                                                 0
        KC_group
                                                 0
        Purebreed_status
                                                 0
        Date of diagnosis_3
                                                 0
                                                 0
        Failuredate
        Date of first suspicion_4
                                                 0
        Date trilostane started_5
                                                14
        Changes to trilostane_6
                                                13
        Stay_vs_stop
                                                 0
        Treated with trilostane_7
                                                 0
        Trilostane starting dose (mg/kg)_8
                                                20
        Trilostane SID/BID_9
                                                13
                                               174
        Censored_10
        Why censored_11
                                               179
        Cause of death_13
                                                42
        Neuro signs
                                                 0
                                                 0
        Complications
        Hypertensive_Yes4
                                                 0
        Oversuppresion?
                                                 0
        Cortisol stayed <250
                                                 0
        Cortisol went <40
                                                 0
        Number_comorbidities
                                                 0
                                                 0
        comorb_UTI
        comorb_dm
                                                 0
        comorb_hypot
                                                 0
        dtype: int64
In [10]: # Creating the categorical_columns
         categorical_columns = [
             'Died',
             'how died',
             'Insurance',
              'Sex',
             'Isneutered',
              'Breed',
              'KC_group',
              'BreedRelativeWeight',
              'Purebreed_status',
              'Changes to trilostane_6',
              'Stay_vs_stop',
              'Treated with trilostane_7',
              'Trilostane SID/BID_9',
              'Censored_10',
              'Why censored_11',
              'Cause of death_13',
```

```
'Neuro signs',
    'Complications',
    'Hypertensive_Yes4',
    'Oversuppresion?',
    'Cortisol stayed <250',
   'Cortisol went <40',
    'Number_comorbidities',
    'comorb_UTI',
    'comorb_dm',
    'comorb_hypot'
]
# Converting the specified columns to categorical (factor)
for col in categorical_columns:
   canine_subset[col] = canine_subset[col].astype('category')
# Generating the descriptive statistics for each categorical variable
descriptive_stats = {}
for col in categorical_columns:
   descriptive_stats[col] = canine_subset[col].value_counts(dropna=False)
# Printing the descriptive statistics
for col, stats in descriptive_stats.items():
   print(f"\nDescriptive statistics for '{col}':")
   print(stats)
```

```
Descriptive statistics for 'Died':
Died
    179
1
0
     40
Name: count, dtype: int64
Descriptive statistics for 'how died':
how died
Euthanasia
           126
             73
Unassisted
              19
Unknown
               1
Name: count, dtype: int64
Descriptive statistics for 'Insurance':
Insurance
Insured
            126
Uninsured
            52
             41
Name: count, dtype: int64
Descriptive statistics for 'Sex':
Sex
male
         115
Female
         104
Name: count, dtype: int64
Descriptive statistics for 'Isneutered':
Isneutered
Yes
    169
       50
Name: count, dtype: int64
Descriptive statistics for 'Breed':
Breed
Crossbreed
47
Terrier - Jack Russell (JRT)
Terrier - Yorkshire (Yorkshire Terrier)
16
Bichon - Frise
13
Terrier - Staffordshire Bull (unspecified) (Staffordshire Bull Terrier (unspecifie
                           13
Terrier - West Highland White (WHWT) (West Highland White Terrier (WHWT))
Retriever - Labrador (Labrador Retriever)
Schnauzer
Spaniel - Cocker (unspecified) (Cocker Spaniel (unspecified))
Terrier - Border (Border Terrier)
Boxer (unspecified)
```

```
Shih-tzu
Dachshund, Miniature
Lhasa Apso
Terrier - Tibetan
Spaniel - Cavalier King Charles (CKCS) (Cavalier King Charles Spaniel (CKCS))
Terrier - Bull (unspecified) (Bull Terrier (unspecified))
Collie - Bearded
Terrier - Scottish
Shepherd Dog - German (Alsatian) (GSD) (unspecified) (German Shepherd Dog (Alsatian)
(GSD) (unspecified))
Dachshund, Standard
Terrier - Boston
Chihuahua, Short-Haired
Terrier - Patterdale
Terrier - Norfolk
Poodle, Standard
Retriever - Golden (Golden Retriever)
Retriever - Flat Coated
Terrier - Australian Silky
Spaniel - Springer, English (English Springer Spaniel)
Terrier - Cairn
Terrier - Fox (unspecified)
Collie - Border
Terrier - Lakeland
Spaniel - King Charles (KCS)
Sheepdog - Shetland (Sheltie)
Mastiff - Bull
Samoyed
Poodle, Miniature
```

```
Bulldog (unspecified)
Whippet (unspecified)
Name: count, dtype: int64
Descriptive statistics for 'KC_group':
KC group
Terrier
           65
Non-KC
           47
           47
Toy
Gundog
           18
Utility
          18
           9
Hound
Pastoral
            9
Working
            6
Name: count, dtype: int64
Descriptive statistics for 'BreedRelativeWeight':
BreedRelativeWeight
1.0
       81
2.0
       73
4.0
    45
3.0
       20
Name: count, dtype: int64
Descriptive statistics for 'Purebreed_status':
Purebreed_status
1.0
     172
2.0
       47
Name: count, dtype: int64
Descriptive statistics for 'Changes to trilostane_6':
Changes to trilostane_6
3.0
       94
 1.0
        55
4.0
       34
 2.0
        23
NaN
        13
Name: count, dtype: int64
Descriptive statistics for 'Stay_vs_stop':
Stay_vs_stop
1.0 172
2.0
       47
Name: count, dtype: int64
Descriptive statistics for 'Treated with trilostane_7':
Treated with trilostane_7
1.0
      206
2.0
       13
Name: count, dtype: int64
Descriptive statistics for 'Trilostane SID/BID_9':
Trilostane SID/BID_9
```

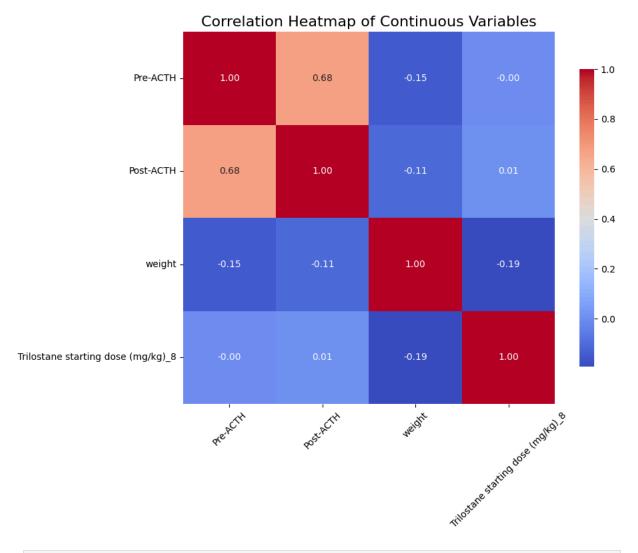
```
SID
           178
BID
            16
NaN
            13
            12
Unknown
Name: count, dtype: int64
Descriptive statistics for 'Censored_10':
Censored_10
NaN
       174
Yes
        40
No
         5
Name: count, dtype: int64
Descriptive statistics for 'Why censored_11':
Why censored 11
NaN
                              179
Moved practice
                               22
No record in last 3 months
                               11
Alive
                                6
Practice cannot contact
                                1
Name: count, dtype: int64
Descriptive statistics for 'Cause of death_13':
Cause of death_13
Unknown
                             55
NaN
                             42
Nervous
                             33
                             24
Digestive
Endocrine (Cushing's)
                             13
Renal/urinary
                             13
Circulatory
                             12
Respiratory
                             10
Musculoskeletal
                              7
Endocrine (not cushing's)
                              5
Immune system/Lymphatic
                              2
Integumental
                              2
Reproductive
Name: count, dtype: int64
Descriptive statistics for 'Neuro signs':
Neuro signs
No
       174
Yes
        45
Name: count, dtype: int64
Descriptive statistics for 'Complications':
Complications
Unknown
           169
            50
Yes
Name: count, dtype: int64
Descriptive statistics for 'Hypertensive_Yes4':
Hypertensive_Yes4
Unknown
           177
Yes
            27
No
            15
```

```
Name: count, dtype: int64
        Descriptive statistics for 'Oversuppresion?':
        Oversuppresion?
        No
             191
        Yes
               28
        Name: count, dtype: int64
        Descriptive statistics for 'Cortisol stayed <250':
        Cortisol stayed <250
        Unknown 119
        Yes
                  68
        No
                   32
        Name: count, dtype: int64
        Descriptive statistics for 'Cortisol went <40':
        Cortisol went <40
        Unknown
                  119
                  79
        No
        Yes
                   21
        Name: count, dtype: int64
        Descriptive statistics for 'Number_comorbidities':
        Number_comorbidities
        1.0
              86
        0.0 78
        2.0 37
        3.0 14
        4.0
            3
        5.0
               1
        Name: count, dtype: int64
        Descriptive statistics for 'comorb_UTI':
        comorb_UTI
        No
             200
        Yes
               19
        Name: count, dtype: int64
        Descriptive statistics for 'comorb_dm':
        comorb_dm
              197
        Nο
        Yes
               22
        Name: count, dtype: int64
        Descriptive statistics for 'comorb_hypot':
        comorb_hypot
              213
        No
        Yes
                6
        Name: count, dtype: int64
In [11]: # Creating the continuous_columns
         continuous_columns = [
            'Pre-ACTH',
             'Post-ACTH',
             'weight',
             'Trilostane starting dose (mg/kg)_8'
```

```
# Generating descriptive statistics for continuous variables
descriptive_stats = canine_subset[continuous_columns].describe()
# Printing the descriptive statistics
print("Descriptive Statistics for Continuous Variables:")
print(descriptive_stats)
# Calculating the correlation matrix
correlation_matrix = canine_subset[continuous_columns].corr()
# Setting up the matplotlib figure
plt.figure(figsize=(10, 8))
# Generating the heatmap with the mask and correct aspect ratio
sns.heatmap(correlation_matrix, annot=True, fmt=".2f", cmap='coolwarm', square=True
# Setting titles and labels
plt.title('Correlation Heatmap of Continuous Variables', fontsize=16)
plt.xticks(rotation=45)
plt.yticks(rotation=0)
plt.tight_layout()
# Showing the plot
plt.show()
```

Descriptive Statistics for Continuous Variables:

| | Pre-ACTH | Post-ACTH | weight | Trilostane starting dose (mg/kg) | 8_(|
|-------|------------|-------------|------------|----------------------------------|-----|
| count | 219.000000 | 219.000000 | 219.000000 | 199.0000 | 900 |
| mean | 53.794064 | 301.045662 | 16.314064 | 3.3015 | 808 |
| std | 101.191170 | 440.164489 | 10.872733 | 1.2756 | 595 |
| min | 0.000000 | 0.000000 | 0.000000 | 1.0000 | 900 |
| 25% | 0.000000 | 0.000000 | 9.200000 | 2.5000 | 900 |
| 50% | 0.000000 | 0.000000 | 12.750000 | 3.0000 | 900 |
| 75% | 88.850000 | 682.500000 | 20.350000 | 4.0000 | 900 |
| max | 508.000000 | 1380.000000 | 65.600000 | 8.0000 | 900 |



```
In [12]: # Converting date columns to datetime format
         date_columns = [
             'BirthDate',
             'Date of first suspicion_4',
             'Date of diagnosis_3',
             'Date trilostane started_5',
             'Failuredate'
         ]
         for col in date_columns:
             canine_subset[col] = pd.to_datetime(canine_subset[col], format='%d-%m-%Y', erro
         # Filtering for uncensored data (Died = 1)
         uncensored_data = canine_subset[canine_subset['Died'] == 1].copy() # Using .copy()
         # Counting of the uncensored entries
         count_uncensored = len(uncensored_data)
         # Earliest and latest dates
         earliest_dates = uncensored_data[date_columns].min()
         latest_dates = uncensored_data[date_columns].max()
         # Date ranges
         date_ranges = latest_dates - earliest_dates
```

```
# Calculating the time intervals using .loc to avoid warnings
uncensored_data.loc[:, 'Time_Birth_to_Suspicion'] = (uncensored_data['Date of first
uncensored_data.loc[:, 'Time_Suspicion_to_Diagnosis'] = (uncensored_data['Date of d
uncensored_data.loc[:, 'Time_Diagnosis_to_Treatment'] = (uncensored_data['Date tril
uncensored_data.loc[:, 'Time_Treatment_to_Failure'] = (uncensored_data['Failuredate
# Mean and median time intervals
mean_intervals = uncensored_data[['Time_Birth_to_Suspicion',
                                   'Time_Suspicion_to_Diagnosis',
                                   'Time_Diagnosis_to_Treatment',
                                   'Time_Treatment_to_Failure']].mean()
median_intervals = uncensored_data[['Time_Birth_to_Suspicion',
                                     'Time_Suspicion_to_Diagnosis',
                                     'Time Diagnosis to Treatment',
                                     'Time_Treatment_to_Failure']].median()
# Printing the results
print(f"Count of Uncensored Entries: {count_uncensored}")
print("\nEarliest Dates:")
print(earliest_dates)
print("\nLatest Dates:")
print(latest_dates)
print("\nDate Ranges:")
print(date_ranges)
print("\nMean Time Intervals (days):")
print(mean_intervals)
print("\nMedian Time Intervals (days):")
print(median_intervals)
```

Earliest Dates:

```
BirthDate
                                    1993-11-11
        Date of first suspicion_4 2010-03-08
                                  2010-03-12
        Date of diagnosis_3
        Date trilostane started_5 2010-03-15
        Failuredate
                                    2010-09-03
        dtype: datetime64[ns]
        Latest Dates:
        BirthDate
                                    2009-02-12
       Date of first suspicion_4 2014-08-21
        Date of diagnosis_3
                                  2014-08-29
        Date trilostane started_5 2014-08-29
        Failuredate
                                    2018-02-27
        dtype: datetime64[ns]
        Date Ranges:
        BirthDate
                                    5572 days
        Date of first suspicion_4 1627 days
       Date of diagnosis_3 1631 days
Date trilostane started_5 1628 days
        Failuredate
                                   2734 days
        dtype: timedelta64[ns]
        Mean Time Intervals (days):
       Time Birth to Suspicion 4020.620112
        Time_Suspicion_to_Diagnosis
                                       45.664804
       Time_Diagnosis_to_Treatment 10.396450
Time_Treatment_to_Failure 503.834320
                                       10.396450
        dtype: float64
        Median Time Intervals (days):
        Time_Birth_to_Suspicion
                                     4024.0
        Time_Suspicion_to_Diagnosis
                                       12.0
                                        0.0
        Time_Diagnosis_to_Treatment
        Time_Treatment_to_Failure
                                      402.0
        dtype: float64
In [13]: # Creating a new column 'Censored' based on 'Censored 10'
         # Using .map() to convert 'Yes' to 1 and 'No' to 0, treating nulls as 0
         canine_subset['Censored'] = canine_subset['Censored_10'].map({'Yes': 1, 'No': 0}).f
         # Printing the updated DataFrame to verify the changes
         print(canine_subset[['Censored_10', 'Censored']].head())
         # Calculating survival time from diagnosis to failure
         canine_subset['Days_Diagnosis_to_Failure'] = (canine_subset['Failuredate'] - canine
         # Printing the updated DataFrame to verify the changes
         print(canine_subset['Days_Diagnosis_to_Failure'].head())
         # Dropping the Censored 10 column
         canine_subset.drop(columns=['Censored_10'], inplace=True)
```

```
# Printing the length of the DataFrame and the remaining column names
 num_rows = len(canine_subset)
 num cols = canine subset.shape[1] # Number of columns
 print(f"Length of DataFrame: {num_rows}")
 print(f"Number of Columns: {num cols}")
 print("Remaining Column Names:", canine_subset.columns.tolist())
 Censored 10 Censored
0
         Yes
         Yes
                      1
1
2
         NaN
                      0
3
         NaN
                      0
4
         NaN
                      0
0
    1778
1
    1842
2
    1260
3
     942
4
     779
Name: Days_Diagnosis_to_Failure, dtype: int64
Length of DataFrame: 219
Number of Columns: 36
Remaining Column Names: ['Died', 'how died', 'Pre-ACTH', 'Post-ACTH', 'Insurance',
'weight', 'BreedRelativeWeight', 'BirthDate', 'Sex', 'Isneutered', 'Breed', 'KC_grou
p', 'Purebreed_status', 'Date of diagnosis_3', 'Failuredate', 'Date of first suspici
on_4', 'Date trilostane started_5', 'Changes to trilostane_6', 'Stay_vs_stop', 'Trea
ted with trilostane_7', 'Trilostane starting dose (mg/kg)_8', 'Trilostane SID/BID_
```

```
In [14]: # Displaying detailed information about the DataFrame
print(canine_subset.info())
print(canine_subset['Died'].head())
```

sis_to_Failure']

9', 'Why censored_11', 'Cause of death_13', 'Neuro signs', 'Complications', 'Hyperte nsive_Yes4', 'Oversuppresion?', 'Cortisol stayed <250', 'Cortisol went <40', 'Number _comorbidities', 'comorb_UTI', 'comorb_dm', 'comorb_hypot', 'Censored', 'Days_Diagno

```
<class 'pandas.core.frame.DataFrame'>
RangeIndex: 219 entries, 0 to 218
Data columns (total 36 columns):
# Column
                                     Non-Null Count Dtype
--- -----
                                     _____
                                     219 non-null
0
   Died
                                                    category
                                     219 non-null category
1
    how died
 2
                                     219 non-null float64
    Pre-ACTH
    Post-ACTH
                                     219 non-null float64
4
                                     219 non-null category
   Insurance
 5
                                     219 non-null float64
    weight
                                    219 non-null category
    BreedRelativeWeight
                                     219 non-null datetime64[ns]
 7
    BirthDate
    Sex
                                     219 non-null category
 9
    Isneutered
                                     219 non-null category
                                     219 non-null category
10 Breed
11 KC_group
                                     219 non-null category
                                     219 non-null
12 Purebreed_status
                                                    category
13 Date of diagnosis 3
                                    219 non-null
                                                    datetime64[ns]
 14 Failuredate
                                     219 non-null
                                                    datetime64[ns]
15 Date of first suspicion_4
                                     219 non-null
                                                    datetime64[ns]
16 Date trilostane started 5
                                     205 non-null
                                                    datetime64[ns]
                                     206 non-null
17 Changes to trilostane_6
                                                    category
18 Stay_vs_stop
                                     219 non-null category
19 Treated with trilostane 7
                                     219 non-null category
 20 Trilostane starting dose (mg/kg)_8 199 non-null
                                                    float64
 21 Trilostane SID/BID_9
                                     206 non-null
                                                    category
 22 Why censored 11
                                     40 non-null
                                                    category
                                     177 non-null category
23 Cause of death_13
 24 Neuro signs
                                     219 non-null category
 25 Complications
                                     219 non-null category
 26 Hypertensive Yes4
                                     219 non-null category
 27 Oversuppresion?
                                    219 non-null category
 28 Cortisol stayed <250
                                    219 non-null category
 29 Cortisol went <40
                                     219 non-null category
 30 Number_comorbidities
                                    219 non-null category
                                     219 non-null
 31 comorb UTI
                                                    category
 32 comorb dm
                                     219 non-null category
33 comorb_hypot
                                     219 non-null
                                                    category
 34 Censored
                                     219 non-null
                                                    int32
35 Days_Diagnosis_to_Failure
                                     219 non-null
                                                    int64
dtypes: category(25), datetime64[ns](5), float64(4), int32(1), int64(1)
memory usage: 28.6 KB
None
0
    0
1
    0
2
    1
3
    1
4
Name: Died, dtype: category
Categories (2, int32): [0, 1]
```

```
In [15]: # The following preprocessing is done for fitting CoxPH and Random Survival Forests
# Calculating Age at diagnosis
canine_subset['Age_at_diagnosis(Years)'] = (canine_subset['Date of diagnosis_3'] -
```

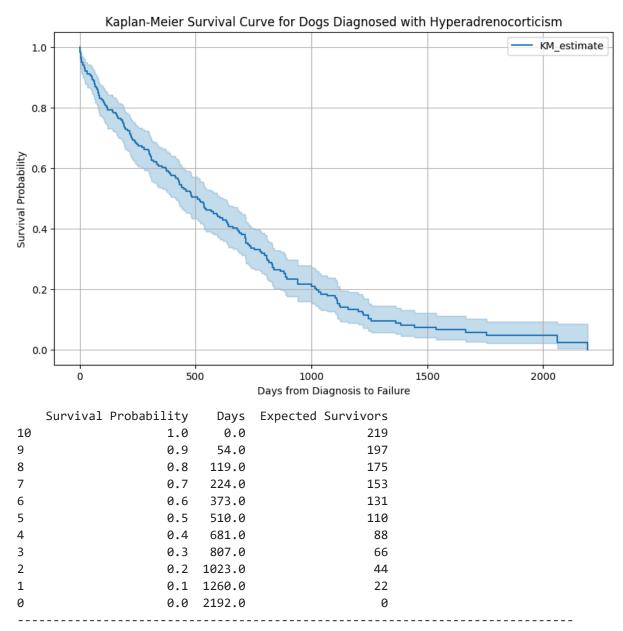
```
# Calculating Time from first suspicion to diagnosis
         canine_subset['Time_suspicion_to_diagnosis(Days)'] = (canine_subset['Date of diagno
         # Rounding the results to 2 decimal places for better readability
         canine_subset['Age_at_diagnosis(Years)'] = canine_subset['Age_at_diagnosis(Years)']
         canine_subset['Time_suspicion_to_diagnosis(Days)'] = canine_subset['Time_suspicion_
         # Displaying the first few rows of the new columns to verify
         print(canine_subset[['Age_at_diagnosis(Years)', 'Time_suspicion_to_diagnosis(Days)'
         # Getting the basic statistics of the new variables
         print("\nStatistics for Age at diagnosis(Years):")
         print(canine_subset['Age_at_diagnosis(Years)'].describe())
         print("\nStatistics for Time from suspicion to diagnosis(Days):")
         print(canine_subset['Time_suspicion_to_diagnosis(Days)'].describe())
           Age_at_diagnosis(Years) Time_suspicion_to_diagnosis(Days)
        0
                             10.04
                                                                    7
                             10.36
                                                                   22
        1
        2
                              8.66
                                                                    6
        3
                             10.79
                                                                   35
        4
                             10.03
                                                                   60
        Statistics for Age at diagnosis(Years):
        count
                219.000000
        mean
                10.991461
                  2.488682
        std
        min
                  4.450000
        25%
                 9.120000
        50%
                  11.010000
        75%
                  12.855000
                  18.000000
        max
        Name: Age_at_diagnosis(Years), dtype: float64
        Statistics for Time from suspicion to diagnosis(Days):
                 219.000000
        count
        mean
                 41.963470
        std
                  81.534866
                  0.000000
        min
        25%
                 6.000000
        50%
                  12.000000
        75%
                  29.500000
                 548.000000
        max
        Name: Time_suspicion_to_diagnosis(Days), dtype: float64
In [16]: # Displaying detailed information about the DataFrame
         print(canine_subset.info())
```

```
<class 'pandas.core.frame.DataFrame'>
       RangeIndex: 219 entries, 0 to 218
       Data columns (total 38 columns):
            Column
                                              Non-Null Count Dtype
       --- -----
                                              _____
        0
            Died
                                              219 non-null
                                                             category
                                              219 non-null category
        1
            how died
                                              219 non-null
            Pre-ACTH
                                                             float64
            Post-ACTH
                                              219 non-null float64
        4
                                              219 non-null
            Insurance
                                                             category
        5
                                              219 non-null float64
            weight
            BreedRelativeWeight
                                              219 non-null category
        7
            BirthDate
                                              219 non-null
                                                             datetime64[ns]
            Sex
                                              219 non-null
                                                            category
            Isneutered
                                              219 non-null
                                                             category
                                              219 non-null category
        10 Breed
        11 KC_group
                                              219 non-null
                                                             category
                                              219 non-null
        12 Purebreed_status
                                                             category
        13 Date of diagnosis 3
                                              219 non-null
                                                             datetime64[ns]
        14 Failuredate
                                              219 non-null
                                                             datetime64[ns]
        15 Date of first suspicion_4
                                              219 non-null
                                                             datetime64[ns]
                                              205 non-null
        16 Date trilostane started 5
                                                             datetime64[ns]
                                              206 non-null
        17 Changes to trilostane_6
                                                             category
                                                            category
        18 Stay_vs_stop
                                              219 non-null
        19 Treated with trilostane 7
                                              219 non-null
                                                             category
        20 Trilostane starting dose (mg/kg)_8 199 non-null
                                                             float64
                                              206 non-null
        21 Trilostane SID/BID_9
                                                             category
        22 Why censored 11
                                              40 non-null
                                                             category
        23 Cause of death_13
                                              177 non-null
                                                             category
        24 Neuro signs
                                              219 non-null category
                                              219 non-null category
        25 Complications
                                              219 non-null
        26 Hypertensive Yes4
                                                             category
        27 Oversuppresion?
                                              219 non-null
                                                             category
        28 Cortisol stayed <250
                                              219 non-null
                                                             category
        29 Cortisol went <40
                                              219 non-null category
        30 Number_comorbidities
                                              219 non-null category
                                              219 non-null
        31 comorb UTI
                                                             category
        32 comorb_dm
                                              219 non-null
                                                             category
        33 comorb_hypot
                                              219 non-null
                                                             category
        34 Censored
                                              219 non-null
                                                             int32
        35 Days_Diagnosis_to_Failure
                                              219 non-null
                                                             int64
        36 Age_at_diagnosis(Years)
                                              219 non-null
                                                             float64
        37 Time_suspicion_to_diagnosis(Days) 219 non-null
                                                             int64
       dtypes: category(25), datetime64[ns](5), float64(5), int32(1), int64(2)
       memory usage: 32.1 KB
       None
In [44]: | #### Preprocessing - 1 ends here
```

Kaplan - Meier Estimator

```
In [18]: # Fitting the Kaplan-Meier estimator
kmf = KaplanMeierFitter()
kmf.fit(durations=canine_subset['Days_Diagnosis_to_Failure'], event_observed=canine
```

```
# Plotting the Kaplan-Meier survival function
plt.figure(figsize=(10, 6))
kmf.plot survival function()
plt.title('Kaplan-Meier Survival Curve for Dogs Diagnosed with Hyperadrenocorticism
plt.xlabel('Days from Diagnosis to Failure')
plt.ylabel('Survival Probability')
plt.grid(True)
plt.show()
# Extracting survival probabilities and time points
survival_function = kmf.survival_function_
time_points = survival_function.index.values
survival_probabilities = survival_function.values.flatten()
# Calculating expected survivors at specific probabilities from 1.0 to 0.0
total_dogs = len(canine_subset) # Total number of dogs
thresholds = [round(i * 0.1, 1) for i in range(11)] # Creating [1.0, 0.9, ..., 0.0
# Initializing a list to store results
results = []
for threshold in thresholds:
   # Finding the closest survival probability to the threshold
   closest_index = (survival_probabilities <= threshold).argmax() # Getting index</pre>
   days_at_threshold = time_points[closest_index]
   expected_survivors = round(total_dogs * threshold) # Rounding the expected sur
   results.append({
        'Survival Probability': threshold,
        'Days': days_at_threshold,
        'Expected Survivors': expected_survivors
   })
# Converting results to a DataFrame for better display
results_df = pd.DataFrame(results)
# Sorting the DataFrame by Survival Probability in descending order
results_df.sort_values(by='Survival Probability', ascending=False, inplace=True)
# Printing the results table
print(results_df)
# Calculating median survival time
median_survival_time = kmf.median_survival_time_
print(f"Median Survival Time for Dogs Diagnosed with Hyperadrenocorticism: {median
```



Median Survival Time for Dogs Diagnosed with Hyperadrenocorticism: 510.00 days

Log Rank Tests

Based on Neutered Status

```
In [19]: # Grouping data based on Isneutered status
group1 = canine_subset[canine_subset['Isneutered'] == 'Yes']
group2 = canine_subset[canine_subset['Isneutered'] == 'No']

# Performing Log-Rank test between the two groups
results_isneutered = logrank_test(group1['Days_Diagnosis_to_Failure'], group2['Days_event_observed_A=group1['Died'], event_observed_B

# Printing the p-value for the comparison
print(f"Log-rank test between Neutered and Non-neutered: p-value = {results_isneute}

# Calculating the Bonferroni-corrected threshold (only one comparison here)
```

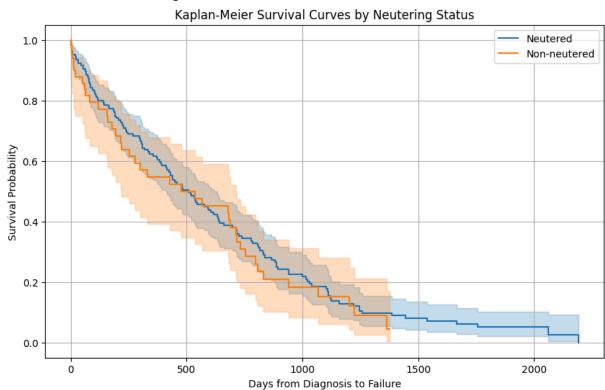
```
alpha = 0.05
n_comparisons = 1 # Since we are only comparing two groups
bonferroni threshold = alpha / n comparisons
print(f"\nBonferroni-corrected significance threshold: {bonferroni_threshold:.4f}")
# Calculating the survival probabilities for both groups
kmf_yes = KaplanMeierFitter()
kmf_no = KaplanMeierFitter()
kmf_yes.fit(group1['Days_Diagnosis_to_Failure'], group1['Died'], label="Neutered")
kmf_no.fit(group2['Days_Diagnosis_to_Failure'], group2['Died'], label="Non-neutered
# Plotting Kaplan-Meier survival curves
plt.figure(figsize=(10, 6))
kmf yes.plot()
kmf_no.plot()
plt.title('Kaplan-Meier Survival Curves by Neutering Status')
plt.xlabel('Days from Diagnosis to Failure')
plt.ylabel('Survival Probability')
plt.grid(True)
plt.legend()
plt.show()
# Defining the thresholds (from 1.0 to 0.0)
thresholds = [round(i * 0.1, 1) for i in range(11)][::-1] # Creates [1.0, 0.9, ...
# Initializing the lists to store results
results_yes = []
results_no = []
for threshold in thresholds:
   # For Neutered group
    closest_index_yes = (kmf_yes.survival_function_.values <= threshold).argmax()</pre>
    days_at_threshold_yes = kmf_yes.survival_function_.index[closest_index_yes]
    expected_survivors_yes = round(len(group1) * threshold)
    results yes.append({
        'Survival Probability': threshold,
        'Days': days_at_threshold_yes,
        'Expected Survivors': expected_survivors_yes
    })
    # For Non-neutered group
    closest_index_no = (kmf_no.survival_function_.values <= threshold).argmax()</pre>
    days_at_threshold_no = kmf_no.survival_function_.index[closest_index_no]
    expected_survivors_no = round(len(group2) * threshold)
    results_no.append({
        'Survival Probability': threshold,
        'Days': days_at_threshold_no,
        'Expected Survivors': expected_survivors_no
    })
# Converting the results to DataFrames for better display
results df yes = pd.DataFrame(results yes)
```

```
results_df_no = pd.DataFrame(results_no)

# Printing the results tables of survival probabilities
print("\nExpected Survival Probabilities for Neutered Dogs:")
print(results_df_yes)
print("\nExpected Survival Probabilities for Non-neutered Dogs:")
print(results_df_no)
```

Log-rank test between Neutered and Non-neutered: p-value = 0.4999

Bonferroni-corrected significance threshold: 0.0500



```
Expected Survival Probabilities for Neutered Dogs:
   Survival Probability Days Expected Survivors
0
                   1.0
                         0.0
                                             169
1
                   0.9
                       63.0
                                             152
2
                   0.8 119.0
                                             135
3
                   0.7
                       240.0
                                             118
4
                   0.6 385.0
                                             101
5
                   0.5 510.0
                                             84
6
                   0.4 643.0
                                             68
7
                   0.3 829.0
                                              51
8
                   0.2 1031.0
                                              34
9
                   0.1 1260.0
                                              17
10
                   0.0 2192.0
                                              0
Expected Survival Probabilities for Non-neutered Dogs:
   Survival Probability Days Expected Survivors
                   1.0
                         0.0
0
                   0.9
                         15.0
1
                                              45
2
                   0.8 82.0
                                              40
3
                   0.7 194.0
                                              35
4
                   0.6 279.0
                                              30
5
                   0.5 536.0
                                              25
6
                   0.4 688.0
                                              20
7
                   0.3 754.0
                                              15
8
                   0.2 940.0
                                              10
                   0.1 1224.0
9
                                              5
                   0.0 0.0
                                              0
```

Based on Purebreed Status

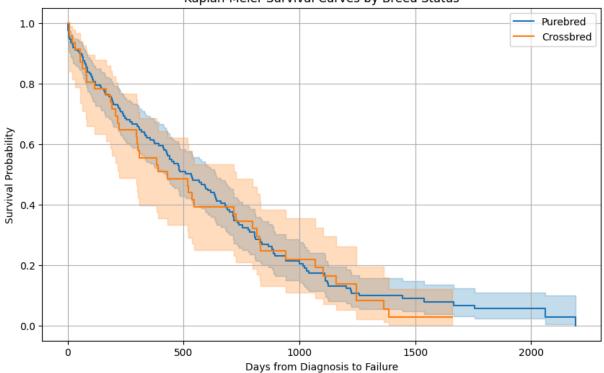
10

```
In [20]: # Grouping data based on 'Purebreed status'
         group1 = canine_subset[canine_subset['Purebreed_status'] == 1] # Purebred
         group2 = canine_subset[canine_subset['Purebreed_status'] == 2] # Crossbred
         # Performing Log-Rank test between the two groups
         results_purebreed = logrank_test(group1['Days_Diagnosis_to_Failure'], group2['Days_
                                          event_observed_A=group1['Died'], event_observed_B=
         # Printing the p-value for the comparison
         print(f"Log-rank test between Purebred and Crossbred: p-value = {results_purebreed.
         # Calculating the Bonferroni-corrected threshold (only one comparison here)
         alpha = 0.05
         n_comparisons = 1 # Since we are only comparing two groups
         bonferroni_threshold = alpha / n_comparisons
         print(f"\nBonferroni-corrected significance threshold: {bonferroni_threshold:.4f}")
         # Calculating the survival probabilities for both groups
         kmf purebred = KaplanMeierFitter()
         kmf_crossbred = KaplanMeierFitter()
         kmf purebred fit(group1['Days Diagnosis to Failure'], group1['Died'], label="Purebr
         kmf_crossbred.fit(group2['Days_Diagnosis_to_Failure'], group2['Died'], label="Cross
         # Plotting Kaplan-Meier survival curves
```

```
plt.figure(figsize=(10, 6))
kmf_purebred.plot()
kmf_crossbred.plot()
plt.title('Kaplan-Meier Survival Curves by Breed Status')
plt.xlabel('Days from Diagnosis to Failure')
plt.ylabel('Survival Probability')
plt.grid(True)
plt.legend()
plt.show()
# Defining the thresholds (from 1.0 to 0.0)
thresholds = [round(i * 0.1, 1) for i in range(11)][::-1] # Creates [1.0, 0.9, ...
# Initializing the lists to store results
results_purebred = []
results_crossbred = []
for threshold in thresholds:
    # For Purebred group
    closest_index_purebred = (kmf_purebred.survival_function_.values <= threshold).</pre>
    days_at_threshold_purebred = kmf_purebred.survival_function_.index[closest_inde
    expected_survivors_purebred = round(len(group1) * threshold)
    results purebred.append({
        'Survival Probability': threshold,
        'Days': days_at_threshold_purebred,
        'Expected Survivors': expected_survivors_purebred
    })
    # For Crossbred group
    closest_index_crossbred = (kmf_crossbred.survival_function_.values <= threshold</pre>
    days_at_threshold_crossbred = kmf_crossbred.survival_function_.index[closest_in
    expected_survivors_crossbred = round(len(group2) * threshold)
    results_crossbred.append({
        'Survival Probability': threshold,
        'Days': days at threshold crossbred,
        'Expected Survivors': expected_survivors_crossbred
    })
# Converting the results to DataFrames for better display
results_df_purebred = pd.DataFrame(results_purebred)
results_df_crossbred = pd.DataFrame(results_crossbred)
# Printing the results tables of survival probabilities
print("\nExpected Survival Probabilities for Purebred Dogs:")
print(results_df_purebred)
print("\nExpected Survival Probabilities for Crossbred Dogs:")
print(results_df_crossbred)
```

Log-rank test between Purebred and Crossbred: p-value = 0.6397

Kaplan-Meier Survival Curves by Breed Status



Expected Survival Probabilities for Purebred Dogs:

| | Survival | Probability | Days | Expected | Survivors |
|----|----------|-------------|--------|----------|-----------|
| 0 | | 1.0 | 0.0 | | 172 |
| 1 | | 0.9 | 50.0 | | 155 |
| 2 | | 0.8 | 119.0 | | 138 |
| 3 | | 0.7 | 237.0 | | 120 |
| 4 | | 0.6 | 395.0 | | 103 |
| 5 | | 0.5 | 533.0 | | 86 |
| 6 | | 0.4 | 681.0 | | 69 |
| 7 | | 0.3 | 807.0 | | 52 |
| 8 | | 0.2 | 1015.0 | | 34 |
| 9 | | 0.1 | 1260.0 | | 17 |
| 10 | | 0.0 | 2192.0 | | 0 |

Expected Survival Probabilities for Crossbred Dogs:

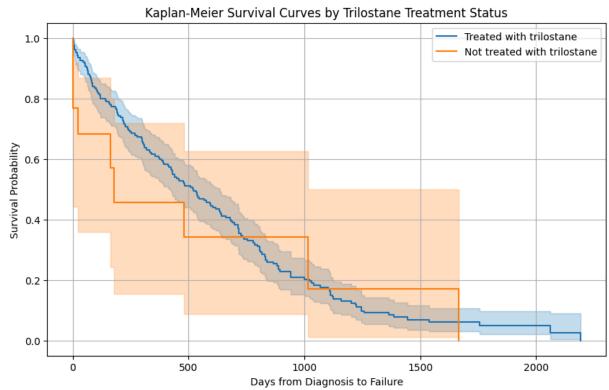
| _ | Apecieu Sulvivai ilobabi | .11(162 | or crossored bogs. |
|---|--------------------------|---------|--------------------|
| | Survival Probability | Days | Expected Survivors |
| 0 | 1.0 | 0.0 | 47 |
| 1 | 0.9 | 54.0 | 42 |
| 2 | 0.8 | 118.0 | 38 |
| 3 | 0.7 | 208.0 | 33 |
| 4 | 0.6 | 304.0 | 28 |
| 5 | 0.5 | 432.0 | 24 |
| 6 | 0.4 | 545.0 | 19 |
| 7 | 0.3 | 817.0 | 14 |
| 8 | 0.2 | 1069.0 | 9 |
| 9 | 0.1 | 1245.0 | 5 |
| 1 | 0.0 | 0.0 | 0 |
| | | | |

Based on treated with trilostane

```
group2 = canine_subset[canine_subset['Treated with trilostane_7'] == 2] # Not trea
# Performing Log-Rank test between the two groups
results_trilostane = logrank_test(group1['Days_Diagnosis_to_Failure'], group2['Days
                                  event_observed_A=group1['Died'], event_observed_B
# Printing the p-value for the comparison
print(f"Log-rank test between Treated and Not Treated with trilostane: p-value = {r
# Calculating the Bonferroni-corrected threshold (only one comparison here)
alpha = 0.05
n_comparisons = 1 # Since we are only comparing two groups
bonferroni_threshold = alpha / n_comparisons
print(f"\nBonferroni-corrected significance threshold: {bonferroni_threshold:.4f}")
# Calculating the survival probabilities for both groups
kmf_treated = KaplanMeierFitter()
kmf_not_treated = KaplanMeierFitter()
kmf_treated.fit(group1['Days_Diagnosis_to_Failure'], group1['Died'], label="Treated
kmf_not_treated.fit(group2['Days_Diagnosis_to_Failure'], group2['Died'], label="Not
# Plotting Kaplan-Meier survival curves
plt.figure(figsize=(10, 6))
kmf treated.plot()
kmf_not_treated.plot()
plt.title('Kaplan-Meier Survival Curves by Trilostane Treatment Status')
plt.xlabel('Days from Diagnosis to Failure')
plt.ylabel('Survival Probability')
plt.grid(True)
plt.legend()
plt.show()
# Defining the thresholds (from 1.0 to 0.0)
thresholds = [round(i * 0.1, 1) for i in range(11)][::-1] # Creates [1.0, 0.9, ...]
# Initializing the lists to store results
results_treated = []
results_not_treated = []
for threshold in thresholds:
   # For Treated group
   closest_index_treated = (kmf_treated.survival_function_.values <= threshold).ar</pre>
   days_at_threshold_treated = kmf_treated.survival_function_.index[closest_index]
   expected_survivors_treated = round(len(group1) * threshold)
   results_treated.append({
        'Survival Probability': threshold,
        'Days': days at threshold treated,
        'Expected Survivors': expected_survivors_treated
   })
   # For Not Treated group
   closest_index_not_treated = (kmf_not_treated.survival_function_.values <= thres</pre>
    days_at_threshold_not_treated = kmf_not_treated.survival_function_.index[closes
```

Log-rank test between Treated and Not Treated with trilostane: p-value = 0.5896

Bonferroni-corrected significance threshold: 0.0500



Expected Survival Probabilities for Dogs Treated with Trilostane:

| | Survival Probability | Days | Expected | Survivors |
|----|----------------------|--------|----------|-----------|
| 0 | 1.0 | 0.0 | | 206 |
| 1 | 0.9 | 63.0 | | 185 |
| 2 | 0.8 | 119.0 | | 165 |
| 3 | 0.7 | 240.0 | | 144 |
| 4 | 0.6 | 376.0 | | 124 |
| 5 | 0.5 | 520.0 | | 103 |
| 6 | 0.4 | 683.0 | | 82 |
| 7 | 0.3 | 807.0 | | 62 |
| 8 | 0.2 | 1023.0 | | 41 |
| 9 | 0.1 | 1245.0 | | 21 |
| 10 | 0.0 | 2192.0 | | 0 |

Expected Survival Probabilities for Dogs Not Treated with Trilostane:

| Survival Probability | Days | Expected Survivors |
|----------------------|---|--|
| 1.0 | 0.0 | 13 |
| 0.9 | 1.0 | 12 |
| 0.8 | 3.0 | 10 |
| 0.7 | 24.0 | 9 |
| 0.6 | 163.0 | 8 |
| 0.5 | 178.0 | 6 |
| 0.4 | 481.0 | 5 |
| 0.3 | 1015.0 | 4 |
| 0.2 | 1015.0 | 3 |
| 0.1 | 1665.0 | 1 |
| 0.0 | 1665.0 | 0 |
| | 1.0 0.9 0.8 0.7 0.6 0.5 0.4 0.3 0.2 | 1.0 0.0 0.9 1.0 0.8 3.0 0.7 24.0 0.6 163.0 0.5 178.0 0.4 481.0 0.3 1015.0 0.2 1015.0 0.1 1665.0 |

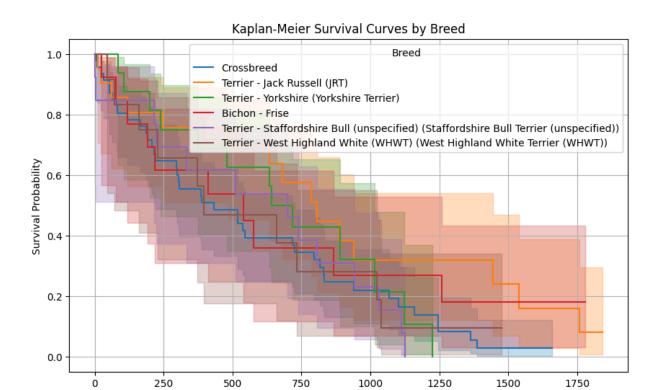
Based on Breeds

```
In [22]: # Counting the number of samples for each breed
breed_counts = canine_subset['Breed'].value_counts()
print(breed_counts)

# Filtering breeds with sufficient representation
sufficient_breeds = breed_counts[breed_counts >= 10].index.tolist()
print("Breeds with sufficient representation:", sufficient_breeds)
```

```
Breed
Crossbreed
47
Terrier - Jack Russell (JRT)
Terrier - Yorkshire (Yorkshire Terrier)
Bichon - Frise
13
Terrier - Staffordshire Bull (unspecified) (Staffordshire Bull Terrier (unspecifie
                           13
Terrier - West Highland White (WHWT) (West Highland White Terrier (WHWT))
Retriever - Labrador (Labrador Retriever)
Schnauzer
Spaniel - Cocker (unspecified) (Cocker Spaniel (unspecified))
Terrier - Border (Border Terrier)
Boxer (unspecified)
Shih-tzu
Dachshund, Miniature
Lhasa Apso
Terrier - Tibetan
Spaniel - Cavalier King Charles (CKCS) (Cavalier King Charles Spaniel (CKCS))
Terrier - Bull (unspecified) (Bull Terrier (unspecified))
Collie - Bearded
Terrier - Scottish
Shepherd Dog - German (Alsatian) (GSD) (unspecified) (German Shepherd Dog (Alsatian)
(GSD) (unspecified))
Dachshund, Standard
Terrier - Boston
Chihuahua, Short-Haired
Terrier - Patterdale
Terrier - Norfolk
Poodle, Standard
Retriever - Golden (Golden Retriever)
Retriever - Flat Coated
```

```
Terrier - Australian Silky
        Spaniel - Springer, English (English Springer Spaniel)
        Terrier - Cairn
        Terrier - Fox (unspecified)
        Collie - Border
        Terrier - Lakeland
        Spaniel - King Charles (KCS)
        Sheepdog - Shetland (Sheltie)
        Mastiff - Bull
        Samoyed
        Poodle, Miniature
        Bulldog (unspecified)
        Whippet (unspecified)
        Name: count, dtype: int64
        Breeds with sufficient representation: ['Crossbreed', 'Terrier - Jack Russell (JR
        T)', 'Terrier - Yorkshire (Yorkshire Terrier)', 'Bichon - Frise', 'Terrier - Staffor
        dshire Bull (unspecified) (Staffordshire Bull Terrier (unspecified))', 'Terrier - We
        st Highland White (WHWT) (West Highland White Terrier (WHWT))']
In [23]: # Creating a dictionary to hold subsets for each breed
         breed_data = {breed: canine_subset[canine_subset['Breed'] == breed] for breed in su
         # Fitting and Plotting Kaplan-Meier curves for each breed
         kmf = KaplanMeierFitter()
         plt.figure(figsize=(10, 6))
         for breed, data in breed_data.items():
             kmf.fit(data['Days_Diagnosis_to_Failure'], event_observed=data['Died'], label=b
             kmf.plot_survival_function()
         plt.title('Kaplan-Meier Survival Curves by Breed')
         plt.xlabel('Time (Days)')
         plt.ylabel('Survival Probability')
         plt.legend(title='Breed')
         plt.grid(True)
         plt.show()
```



Time (Days)

```
In [24]: # Performing Log-rank tests between pairs of breeds
breeds_to_compare = list(sufficient_breeds) # List of breeds to compare
for i in range(len(breeds_to_compare)):
    for j in range(i + 1, len(breeds_to_compare)):
        group1 = breed_data[breeds_to_compare[i]]
        group2 = breed_data[breeds_to_compare[j]]

    results = logrank_test(
        group1['Days_Diagnosis_to_Failure'],
        group2['Days_Diagnosis_to_Failure'],
        event_observed_A=group1['Died'],
        event_observed_B=group2['Died']
)

    print(f"Log-Rank Test between {breeds_to_compare[i]} and {breeds_to_compare
```

```
Log-Rank Test between Crossbreed and Terrier - Jack Russell (JRT): p-value = 0.03642
641759435863
Log-Rank Test between Crossbreed and Terrier - Yorkshire (Yorkshire Terrier): p-valu
e = 0.6692473130343376
Log-Rank Test between Crossbreed and Bichon - Frise: p-value = 0.36841829323395925
Log-Rank Test between Crossbreed and Terrier - Staffordshire Bull (unspecified) (Sta
ffordshire Bull Terrier (unspecified)): p-value = 0.7709627160434211
Log-Rank Test between Crossbreed and Terrier - West Highland White (WHWT) (West High
land White Terrier (WHWT)): p-value = 0.8337970167715292
Log-Rank Test between Terrier - Jack Russell (JRT) and Terrier - Yorkshire (Yorkshir
e Terrier): p-value = 0.2577114515491709
Log-Rank Test between Terrier - Jack Russell (JRT) and Bichon - Frise: p-value = 0.5
158772885275844
Log-Rank Test between Terrier - Jack Russell (JRT) and Terrier - Staffordshire Bull
(unspecified) (Staffordshire Bull Terrier (unspecified)): p-value = 0.12143454232807
Log-Rank Test between Terrier - Jack Russell (JRT) and Terrier - West Highland White
(WHWT) (West Highland White Terrier (WHWT)): p-value = 0.24664745778391245
Log-Rank Test between Terrier - Yorkshire (Yorkshire Terrier) and Bichon - Frise: p-
value = 0.87309074220082
Log-Rank Test between Terrier - Yorkshire (Yorkshire Terrier) and Terrier - Stafford
shire Bull (unspecified) (Staffordshire Bull Terrier (unspecified)): p-value = 0.539
480393516573
Log-Rank Test between Terrier - Yorkshire (Yorkshire Terrier) and Terrier - West Hig
hland White (WHWT) (West Highland White Terrier (WHWT)): p-value = 0.740040465691326
Log-Rank Test between Bichon - Frise and Terrier - Staffordshire Bull (unspecified)
(Staffordshire Bull Terrier (unspecified)): p-value = 0.4652353594419921
Log-Rank Test between Bichon - Frise and Terrier - West Highland White (WHWT) (West
Highland White Terrier (WHWT)): p-value = 0.7182020624984118
Log-Rank Test between Terrier - Staffordshire Bull (unspecified) (Staffordshire Bull
Terrier (unspecified)) and Terrier - West Highland White (WHWT) (West Highland White
Terrier (WHWT)): p-value = 0.9723712953533593
```

Preprocessing - Part 2 : For CoxPH and Random Survival Forests

```
print(f"Total Entries: {len(column)}")
    print(f"Null Count: {null_count}")
   print(tabulate(summary df, headers='keys', tablefmt='pretty', showindex=False))
# Continuous columns summary function
def continuous_column_summary(df, column_name):
   column = df[column_name]
   null_count = column.isnull().sum()
   summary_data = {
        'Statistic': ['Null Count', 'Total Entries', 'Mean', 'Median', 'Min', 'Max'
        'Value': [
            null_count,
            len(column),
            round(column.mean(), 2),
            round(column.median(), 2),
            round(column.min(), 2),
            round(column.max(), 2),
            round(column.std(), 2)
        ]
   }
   summary_df = pd.DataFrame(summary_data)
   print(f"\nSummary for Continuous Column: {column_name}")
   print(tabulate(summary_df, headers='keys', tablefmt='pretty', showindex=False))
# Categorical columns
cat_columns = [
    'Died','Neuro signs', 'Complications', 'Hypertensive_Yes4', 'Oversuppresion?',
    'Cortisol stayed <250', 'Cortisol went <40', 'Number_comorbidities',
    'comorb_UTI', 'comorb_dm', 'comorb_hypot', 'Sex', 'Isneutered',
    'BreedRelativeWeight', 'Purebreed_status', 'Changes to trilostane_6',
    'Stay_vs_stop', 'Treated with trilostane_7', 'Trilostane SID/BID_9'
]
# Continuous columns
cont columns = [
    'Days_Diagnosis_to_Failure', 'Pre-ACTH', 'Post-ACTH', 'weight', 'Trilostane sta
    'Age_at_diagnosis(Years)', 'Time_suspicion_to_diagnosis(Days)'
]
# Generating the summaries for categorical columns
print("CATEGORICAL COLUMNS SUMMARY")
print("=" * 50)
for column in cat_columns:
   categorical_column_summary(canine_subset, column)
# Generating the summaries for continuous columns
print("\n\nCONTINUOUS COLUMNS SUMMARY")
print("=" * 50)
for column in cont_columns:
   continuous_column_summary(canine_subset, column)
```

Summary for Categorical Column: Died

Total Entries: 219
Null Count: 0

| + | | ++ | + |
|------|---------------------|-------|------------|
| Unio | ue Values | Count | Percentage |
| + | . – – – – – – – – – | .++ | + |
| 1 | 0.0 | 40.0 | 18.26 |
| İ | 1.0 | 179.0 | 81.74 |
| + | | ++ | + |

Summary for Categorical Column: Neuro signs

Total Entries: 219

Null Count: 0

| + | • | • | • |
|--------|-------------|------------------|---|
| No Yes | 174 45 | 79.45 20.55 | |

Summary for Categorical Column: Complications

Total Entries: 219

Null Count: 0

| Unique Values | | Percentage | • |
|---------------|---------------|----------------|---|
| Unknown Yes | 169 50 | 77.17 22.83 | |

Summary for Categorical Column: Hypertensive_Yes4

Total Entries: 219

Null Count: 0

| Unique Values | + Count + | ++ Percentage ++ |
|----------------|---------------------|--------------------------------|
| Unknown Yes No | 177 27 15 | 80.82 12.33 6.85 |

Summary for Categorical Column: Oversuppresion?

Total Entries: 219

Null Count: 0

| + Unique Values | • | • | • |
|----------------------|---|-------------------------|---------|
| + Yes No | | + 12.79 87.21 | + + |

Summary for Categorical Column: Cortisol stayed <250

Total Entries: 219

Null Count: 0

| Unique Values Coun | t Percentage |
|----------------------|----------------|
| Yes 68 | 31.05 |
| Unknown 119 | 54.34 |
| No 32 | 14.61 |

Summary for Categorical Column: Cortisol went <40

Total Entries: 219

Null Count: 0

| Unique Values | + Count + | ++ Percentage |
|---------------|-------------------|--------------------|
| Yes | 21 | 9.59 |
| Unknown | 119 | 54.34 |
| No | 79 | 36.07 |

Summary for Categorical Column: Number_comorbidities

Total Entries: 219
Null Count: 0

| + Unique Values | + Count | H+ Percentage |
|------------------|--------------|--------------------|
| + | + | |
| 0.0 | 78.0 | 35.62 |
| 1.0 | 86.0 | 39.27 |
| 3.0 | 14.0 | 6.39 |
| 2.0 | 37.0 | 16.89 |
| 4.0 | 3.0 | 1.37 |
| 5.0 | 1.0 | 0.46 |
| + | + | ++ |

Summary for Categorical Column: comorb_UTI

Total Entries: 219

Null Count: 0

| + | Count | Percentage | İ |
|--------|-------|------------|---|
| No Yes | 200 | 91.32 | |
| | 19 | 8.68 | |

Summary for Categorical Column: comorb_dm

Total Entries: 219

Null Count: 0

+-----+
| Unique Values | Count | Percentage |
+-----+
| No | 197 | 89.95 |
| Yes | 22 | 10.05 |

Summary for Categorical Column: comorb_hypot

Total Entries: 219

Null Count: 0

| Unique Values | Count | |
|---------------|----------|-------|
| No Yes | 213 6 | 97.26 |

Summary for Categorical Column: Sex

Total Entries: 219

Null Count: 0

| ++ | + | + |
|---------------|-------|------------|
| Unique Values | Count | Percentage |
| ++ | + | + |
| male | 115 | 52.51 |
| Female | 104 | 47.49 |
| ++ | + | + |

Summary for Categorical Column: Isneutered

Total Entries: 219

Null Count: 0

| + Unique Values | | • |
|----------------------|-----|-------|
| + | ++ | + |
| Yes | 169 | 77.17 |
| No | 50 | 22.83 |
| + | ++ | + |

Summary for Categorical Column: BreedRelativeWeight

Total Entries: 219

Null Count: 0

| + | + | ++ |
|---------------|-------|------------|
| Unique Values | Count | Percentage |
| + | + | ·+ |
| 1.0 | 81.0 | 36.99 |
| 2.0 | 73.0 | 33.33 |
| 4.0 | 45.0 | 20.55 |
| 3.0 | 20.0 | 9.13 |
| + | + | ·+ |

Summary for Categorical Column: Purebreed_status

Total Entries: 219

Null Count: 0

| 1.0 172.0 78.54 2.0 47.0 21.46 | + Unique Valu | • | • | • |
|---|--------------------|---|---|---|
| | | | : | |

Summary for Categorical Column: Changes to trilostane_6

Total Entries: 219

Null Count: 13

| + | | ++ |
|---------------|---------|------------|
| Unique Values | Count | Percentage |
| + | | ++ |

| 1.0 | 55.0 | 25.11 | |
|-----|------|-------|--|
| 2.0 | 23.0 | 10.5 | |
| 4.0 | 34.0 | 15.53 | |
| nan | 0.0 | 0.0 | |
| 3.0 | 94.0 | 42.92 | |
| | | | |

+----+

Summary for Categorical Column: Stay_vs_stop

Total Entries: 219 Null Count: 0

| + | Count | • |
|-----|-------------------|-------|
| 1.0 | 172.0 47.0 | 78.54 |

Summary for Categorical Column: Treated with trilostane_7

Total Entries: 219

Null Count: 0

| ++ Unique Values Count Percentage | | |
|---|---------|-------|
| + | | ++ |
| 1.0 | 206.0 | 94.06 |
| 2.0 | 13.0 | 5.94 |
| + | · | ++ |

Summary for Categorical Column: Trilostane SID/BID_9

Total Entries: 219 Null Count: 13

| + | | |
|---------------|-----|-------|
| Unique Values | • | |
| SID | 178 | 81.28 |
| BID | 16 | 7.31 |
| Unknown | 12 | 5.48 |
| nan | 0 | 0.0 |
| | | |

+----+

CONTINUOUS COLUMNS SUMMARY

Summary for Continuous Column: Days_Diagnosis_to_Failure

| + | + |
|--------------------|--------|
| Statistic | Value |
| + | ++ |
| Null Count | 0.0 |
| Total Entries | 219.0 |
| Mean | 517.79 |
| Median | 424.0 |
| Min | 1.0 |
| Max | 2192.0 |
| Standard Deviation | 465.73 |
| + | ++ |
| | |

Summary for Continuous Column: Pre-ACTH

| Τ. | | |
|----|--------------------|--------|
| | Statistic | Value |
| İ | Null Count | 0.0 |
| | Total Entries | 219.0 |
| | Mean | 53.79 |
| | Median | 0.0 |
| | Min | 0.0 |
| | Max | 508.0 |
| | Standard Deviation | 101.19 |
| | | |

+----+

Summary for Continuous Column: Post-ACTH

| + | | + | L |
|---|--------------------|--------|---|
| | Statistic | Value | |
| i | Null Count | 0.0 | |
| | Total Entries | 219.0 | |
| | Mean | 301.05 | |
| | Median | 0.0 | |
| | Min | 0.0 | |
| | Max | 1380.0 | |
| | Standard Deviation | 440.16 | |
| | | | |

+----+

Summary for Continuous Column: weight

| L | |
|--------------------|-------|
| Statistic | Value |
| Null Count | 0.0 |
| Total Entries | 219.0 |
| Mean | 16.31 |
| Median | 12.75 |
| Min | 0.0 |
| Max | 65.6 |
| Standard Deviation | 10.87 |

+-----

Summary for Continuous Column: Trilostane starting dose (mg/kg)_8

| İ | Statistic | Value |
|---|--------------------|-------|
| | Null Count | 20.0 |
| | Total Entries | 219.0 |
| | Mean | 3.3 |
| | Median | 3.0 |
| | Min | 1.0 |
| | Max | 8.0 |
| | Standard Deviation | 1.28 |
| | | |

+----+

Summary for Continuous Column: Age_at_diagnosis(Years)

| + | ++ | | | |
|-----------|-------|--|--|--|
| Statistic | Value | | | |
| + | | | | |

```
Null Count | 0.0 |
  Total Entries | 219.0 |
     Mean
              | 10.99 |
     Median
              | 11.01 |
     Min
              4.45
      Max | 18.0 |
| Standard Deviation | 2.49 |
+----+
Summary for Continuous Column: Time_suspicion_to_diagnosis(Days)
+----+
| Statistic | Value |
+----+
   Null Count | 0.0 |
 Total Entries | 219.0 |
    Mean | 41.96 |
Median | 12.0 |
    Min | 0.0 |
Max | 548.0 |
| Standard Deviation | 81.53 |
+----+
```

```
In [26]: # Creating a list of columns to include in cox_data
         columns_to_include = [
             # Clinical factors
             'Pre-ACTH', 'Post-ACTH', 'weight',
             'Neuro signs', 'Complications', 'Hypertensive_Yes4', 'Oversuppresion?',
             'Cortisol stayed <250', 'Cortisol went <40', 'Number_comorbidities',
             'comorb_UTI', 'comorb_dm', 'comorb_hypot',
             # Demographic factors
             'Sex', 'Isneutered', 'BreedRelativeWeight', 'Purebreed_status',
             # Treatment-related factors
             'Stay_vs_stop', 'Treated with trilostane_7',
             # Time-related factors
             'Age_at_diagnosis(Years)', 'Time_suspicion_to_diagnosis(Days)',
             # Outcome and event indicator
             'Days_Diagnosis_to_Failure', 'Died'
         # Copying the selected columns to cox_data
         cox_data = canine_subset[columns_to_include].copy()
         # Displaying the first few rows of cox_data to verify
         print(cox_data.head())
         # Print the column names of cox_data one by one
         print("Column names in cox_data:")
         for col in cox_data.columns:
             print(col)
         # Print the number of columns
         num_cols = len(cox_data.columns)
```

```
print("\nTotal number of columns in cox_data:", num_cols)

# Get the number of rows in cox_subset
num_rows = len(cox_data)
print("Number of rows in cox_subset:", num_rows)
```

```
Pre-ACTH Post-ACTH weight Neuro signs Complications Hypertensive_Yes4 \
       0.0 0.0
                         7.55
0
                                      No
                                               Unknown
       0.0
                 0.0
                         7.20
1
                                       No
                                               Unknown
                                                                 Unknown
2
     267.0
                                                                 Unknown
                922.0 12.50
                                       No
                                               Unknown
3
       0.0
                0.0 19.00
                                       No
                                               Unknown
                                                                 Unknown
4
     145.0
                684.0 34.25
                                       No
                                                   Yes
                                                                     Yes
 Oversuppresion? Cortisol stayed <250 Cortisol went <40 Number_comorbidities \
             Yes
                                 Yes
                                                   Yes
                                                                        0.0
0
             Yes
                              Unknown
                                               Unknown
                                                                        1.0
1
2
             Yes
                                 Yes
                                                                        3.0
                                                    No
3
             Yes
                                  No
                                                    No
                                                                        0.0
4
             Yes
                                  Yes
                                                   Yes
                                                                        1.0
          Sex Isneutered BreedRelativeWeight Purebreed status Stay vs stop \
                                        1.0
         male
                     Yes
                                                         1.0
                                                                      1.0
0
  . . .
  ... Female
                     Yes
                                        1.0
                                                         1.0
                                                                      1.0
1
                     Yes
                                        2.0
                                                         1.0
                                                                      1.0
2
         male
  . . .
3 ... Female
                     Yes
                                        1.0
                                                         1.0
                                                                      2.0
4
         male
                     Yes
                                        2.0
                                                         1.0
                                                                      2.0
 Treated with trilostane_7 Age_at_diagnosis(Years) \
0
                       1.0
                                             10.04
                       1.0
                                             10.36
1
2
                       1.0
                                             8.66
3
                       1.0
                                            10.79
4
                       1.0
                                            10.03
 Time_suspicion_to_diagnosis(Days) Days_Diagnosis_to_Failure Died
                                                       1778
                                22
1
                                                       1842
                                                                0
2
                                 6
                                                       1260
                                                                1
3
                                35
                                                        942
                                                                1
4
                                60
                                                        779
                                                                1
[5 rows x 23 columns]
Column names in cox_data:
Pre-ACTH
Post-ACTH
weight
Neuro signs
Complications
Hypertensive_Yes4
Oversuppresion?
Cortisol stayed <250
Cortisol went <40
Number_comorbidities
comorb_UTI
comorb_dm
comorb_hypot
Sex
Isneutered
BreedRelativeWeight
Purebreed_status
Stay_vs_stop
Treated with trilostane_7
```

```
Age_at_diagnosis(Years)
        Time_suspicion_to_diagnosis(Days)
        Days Diagnosis to Failure
        Died
        Total number of columns in cox data: 23
        Number of rows in cox subset: 219
In [27]: # Encoding Sex: Male as 0 and Female as 1
         cox_data['Sex'] = (cox_data['Sex'].str.lower() == 'female').astype(int)
         # Displaying the unique values and their counts for the new encoded columns
         print("Encoded Sex column:")
         print(cox_data['Sex'].value_counts(dropna=False))
         # Checking for remaining null values
         print("\nNull values in encoded columns:")
         print(cox_data[['Sex']].isnull().sum())
         # Get the number of rows in cox_subset
         num_rows = len(cox_data)
         print("Number of rows in cox_subset:", num_rows)
        Encoded Sex column:
        Sex
        0
             115
             104
        1
        Name: count, dtype: int64
        Null values in encoded columns:
        Sex
        dtype: int64
        Number of rows in cox_subset: 219
In [28]: # Converting specified columns from float to int, keeping NaNs as is
         cox_data['Died'] = cox_data['Died'].astype('Int64')
         print("Died unique values:", cox_data['Died'].unique())
         cox_data['Number_comorbidities'] = cox_data['Number_comorbidities'].astype('Int64')
         print("Number_comorbidities unique values:", cox_data['Number_comorbidities'].uniqu
         cox_data['BreedRelativeWeight'] = cox_data['BreedRelativeWeight'].astype('Int64')
         print("BreedRelativeWeight unique values:", cox_data['BreedRelativeWeight'].unique(
         cox_data['Purebreed_status'] = cox_data['Purebreed_status'].astype('Int64')
         print("Purebreed_status unique values:", cox_data['Purebreed_status'].unique())
         cox_data['Stay_vs_stop'] = cox_data['Stay_vs_stop'].astype('Int64')
         print("Stay_vs_stop unique values:", cox_data['Stay_vs_stop'].unique())
         cox_data['Treated with trilostane_7'] = cox_data['Treated with trilostane_7'].astyp
         print("Treated with trilostane_7 unique values:", cox_data['Treated with trilostane
         # Get the number of rows in cox_subset
         num_rows = len(cox_data)
         print("Number of rows in cox_subset:", num_rows)
```

```
Died unique values: <IntegerArray>
        Length: 2, dtype: Int64
        Number_comorbidities unique values: <IntegerArray>
        [0, 1, 3, 2, 4, 5]
        Length: 6, dtype: Int64
        BreedRelativeWeight unique values: <IntegerArray>
        [1, 2, 4, 3]
        Length: 4, dtype: Int64
        Purebreed_status unique values: <IntegerArray>
        [1, 2]
        Length: 2, dtype: Int64
        Stay vs stop unique values: <IntegerArray>
        [1, 2]
        Length: 2, dtype: Int64
        Treated with trilostane_7 unique values: <IntegerArray>
        Length: 2, dtype: Int64
        Number of rows in cox subset: 219
In [29]: # Suppressing specific warnings
         warnings.filterwarnings("ignore", category=FutureWarning)
         # Encoding Neuro_signs: No as 0, Yes as 1
         cox data['Neuro signs'] = cox data['Neuro signs'].replace({'Yes': 1, 'No': 0})
         print("Neuro_signs unique values:", cox_data['Neuro signs'].unique())
         # Encoding Complications: No as 0, Yes as 1
         cox data['Complications'] = cox data['Complications'].replace({'Yes': 1, 'Unknown':
         print("Complications unique values:", cox_data['Complications'].unique())
         # Encoding Hypertensive Yes4: No as 0, Yes as 1
         cox_data['Hypertensive_Yes4'] = cox_data['Hypertensive_Yes4'].replace({'Yes': 1, 'N
         print("Hypertensive_Yes4 unique values:", cox_data['Hypertensive_Yes4'].unique())
         # Encoding Oversuppresion: No as 0, Yes as 1
         cox_data['Oversuppresion?'] = cox_data['Oversuppresion?'].replace({'Yes': 1, 'No':
         print("Oversuppresion unique values:", cox_data['Oversuppresion?'].unique())
         # Encoding Cortisol stayed <250: No as 0, Yes as 1
         cox data['Cortisol stayed <250'] = cox data['Cortisol stayed <250'].replace({'Yes':</pre>
         print("Cortisol stayed <250 unique values:", cox_data['Cortisol stayed <250'].uniqu</pre>
         # Encoding Cortisol went <40: No as 0, Yes as 1
         cox_data['Cortisol went <40'] = cox_data['Cortisol went <40'].replace({'Yes': 1, 'N</pre>
         print("Cortisol went <40 unique values:", cox_data['Cortisol went <40'].unique())</pre>
         # Encoding comorb UTI: No as 0, Yes as 1
         cox_data['comorb_UTI'] = cox_data['comorb_UTI'].replace({'Yes': 1, 'No': 0})
         print("comorb UTI unique values:", cox data['comorb UTI'].unique())
         # Encoding comorb_dm: No as 0, Yes as 1
         cox data['comorb dm'] = cox data['comorb dm'].replace({'Yes': 1, 'No': 0})
         print("comorb_dm unique values:", cox_data['comorb_dm'].unique())
         # Encoding comorb_hypot: No as 0, Yes as 1
```

```
cox_data['comorb_hypot'] = cox_data['comorb_hypot'].replace({'Yes': 1, 'No': 0})
         print("comorb_hypot unique values:", cox_data['comorb_hypot'].unique())
         # Encoding Isneutered: No as 0, Yes as 1
         cox_data['Isneutered'] = cox_data['Isneutered'].replace({'Yes': 1, 'No': 0})
         print("Isneutered unique values:", cox_data['Isneutered'].unique())
         # Get the number of rows in cox_subset
         num rows = len(cox data)
         print("Number of rows in cox_subset:", num_rows)
         # 'Changes to trilostane_6','Trilostane SID/BID_9' 'Trilostane starting dose (mg/kg
        Neuro_signs unique values: [0, 1]
        Categories (2, int64): [0, 1]
        Complications unique values: [2, 1]
        Categories (2, int64): [2, 1]
        Hypertensive_Yes4 unique values: [2, 1, 0]
        Categories (3, int64): [0, 2, 1]
        Oversuppresion unique values: [1, 0]
        Categories (2, int64): [0, 1]
        Cortisol stayed <250 unique values: [1, 2, 0]
        Categories (3, int64): [0, 2, 1]
        Cortisol went <40 unique values: [1, 2, 0]
        Categories (3, int64): [0, 2, 1]
        comorb_UTI unique values: [0, 1]
        Categories (2, int64): [0, 1]
        comorb dm unique values: [0, 1]
        Categories (2, int64): [0, 1]
        comorb_hypot unique values: [0, 1]
        Categories (2, int64): [0, 1]
        Isneutered unique values: [1, 0]
        Categories (2, int64): [0, 1]
        Number of rows in cox_subset: 219
In [30]: # Printing the column names of cox_data
         print("Column names in cox data:")
         for col in cox_data.columns:
             print(col)
         # Printing the number of columns
         num_cols = len(cox_data.columns)
         print("\nTotal number of columns in cox_data:", num_cols)
         print(cox_data.head(2))
         # Getting the number of rows in cox_subset
         num rows = len(cox data)
         print("Number of rows in cox_subset:", num_rows)
```

```
Column names in cox_data:
Pre-ACTH
Post-ACTH
weight
Neuro signs
Complications
Hypertensive_Yes4
Oversuppresion?
Cortisol stayed <250
Cortisol went <40
Number_comorbidities
comorb_UTI
comorb_dm
comorb_hypot
Sex
Isneutered
BreedRelativeWeight
Purebreed_status
Stay_vs_stop
Treated with trilostane_7
Age_at_diagnosis(Years)
Time_suspicion_to_diagnosis(Days)
Days_Diagnosis_to_Failure
Died
Total number of columns in cox_data: 23
   Pre-ACTH Post-ACTH weight Neuro signs Complications Hypertensive_Yes4 \
        0.0
                   0.0
                          7.55
                                         0
                                                       2
0
                  0.0
                          7.20
1
        0.0
                                         0
                                                       2
                                                                         2
 Oversuppresion? Cortisol stayed <250 Cortisol went <40 \
0
                1
                                     1
                1
                                     2
                                                       2
1
  Number_comorbidities ... Sex Isneutered BreedRelativeWeight \
0
                                          1
                      0 ...
                               0
                                                              1
1
                        . . .
                               1
                                                              1
  Purebreed_status Stay_vs_stop Treated with trilostane_7 \
0
                 1
                               1
                  1
1
                               1
                                                          1
  Age_at_diagnosis(Years) Time_suspicion_to_diagnosis(Days) \
0
                                                            7
                     10.04
1
                     10.36
                                                           22
  Days_Diagnosis_to_Failure Died
0
                        1778
                        1842
1
[2 rows x 23 columns]
Number of rows in cox_subset: 219
```

Subset: final_cox_data is used for CoxPH Models

```
In [31]: # Preparing the DataFrame for fitting CoxPH Model
         final_cox_data = cox_data[['Days_Diagnosis_to_Failure',
                 'Died','Neuro signs', 'Complications', 'Hypertensive_Yes4', 'Oversuppresion
                 'Cortisol stayed <250', 'Cortisol went <40', 'Number_comorbidities',
                 'comorb_UTI', 'comorb_dm', 'comorb_hypot', 'Sex', 'Isneutered',
                 'BreedRelativeWeight', 'Purebreed_status',
                 'Stay_vs_stop', 'Treated with trilostane_7', 'Pre-ACTH', 'Post-ACTH', 'weig
                 'Age_at_diagnosis(Years)', 'Time_suspicion_to_diagnosis(Days)']]
         # Getting the number of rows in cox_subset
         num_rows = len(final_cox_data)
         print("Number of rows in cox_subset:", num_rows)
         # Creating an instance of the CoxPHFitter
         cph = CoxPHFitter()
         cph.fit(final_cox_data, duration_col='Days_Diagnosis_to_Failure', event_col='Died')
         # Printing the summary of the model
         cph.print_summary()
```

Number of rows in cox_subset: 219

| model | lifelines.CoxPHFitter | | | | |
|---------------------------|-----------------------------|--|--|--|--|
| duration col | 'Days_Diagnosis_to_Failure' | | | | |
| event col | 'Died' | | | | |
| baseline estimation | breslow | | | | |
| number of observations | 219 | | | | |
| number of events observed | 179 | | | | |
| partial log-likelihood | -766.83 | | | | |
| time fit was run | 2024-11-23 18:25:33 UTC | | | | |

| | coef | exp(coef) | se(coef) | coef lower 95% | coef upper 95% | exp(coef) lower 95% | ut exp(c |
|-----------------------------------|-------|-----------|----------|----------------------|----------------------|---------------------------|-------------|
| Neuro signs | 0.49 | 1.64 | 0.20 | 0.10 | 0.89 | 1.10 | |
| Complications | -0.14 | 0.87 | 0.30 | -0.72 | 0.44 | 0.49 | |
| Hypertensive_Yes4 | -0.13 | 0.88 | 0.16 | -0.44 | 0.19 | 0.64 | |
| Oversuppresion? | -0.42 | 0.65 | 0.24 | -0.90 | 0.05 | 0.41 | |
| Cortisol stayed <250 | 0.37 | 1.45 | 0.23 | -0.09 | 0.83 | 0.91 | |
| Cortisol went <40 | -0.01 | 0.99 | 0.18 | -0.36 | 0.34 | 0.70 | |
| Number_comorbidities | -0.19 | 0.82 | 0.10 | -0.39 | 0.01 | 0.68 | |
| comorb_UTI | 0.92 | 2.50 | 0.28 | 0.37 | 1.46 | 1.45 | |
| comorb_dm | 0.21 | 1.23 | 0.33 | -0.45 | 0.87 | 0.64 | |
| comorb_hypot | -0.39 | 0.67 | 0.54 | -1.45 | 0.66 | 0.23 | |
| Sex | -0.06 | 0.94 | 0.17 | -0.40 | 0.28 | 0.67 | |
| Isneutered | -0.11 | 0.89 | 0.20 | -0.50 | 0.27 | 0.61 | |
| BreedRelativeWeight | -0.11 | 0.89 | 0.13 | -0.36 | 0.14 | 0.70 | |
| Purebreed_status | 0.20 | 1.22 | 0.33 | -0.45 | 0.85 | 0.64 | |
| Stay_vs_stop | 0.16 | 1.18 | 0.22 | -0.27 | 0.59 | 0.76 | |
| Treated with trilostane_7 | -0.35 | 0.70 | 0.46 | -1.26 | 0.55 | 0.28 | |
| Pre-ACTH | 0.00 | 1.00 | 0.00 | -0.00 | 0.00 | 1.00 | |
| Post-ACTH | -0.00 | 1.00 | 0.00 | -0.00 | 0.00 | 1.00 | |
| weight | 0.01 | 1.01 | 0.01 | -0.00 | 0.03 | 1.00 | |
| Age_at_diagnosis(Years) | 0.16 | 1.18 | 0.04 | 0.09 | 0.24 | 1.09 | |
| Time_suspicion_to_diagnosis(Days) | -0.00 | 1.00 | 0.00 | -0.00 | 0.00 | 1.00 | |

Partial AIC 1575.65
log-likelihood ratio test 53.45 on 21 df
-log2(p) of II-ratio test 13.04

```
In [32]: mpl.rcParams['figure.max_open_warning'] = 50
# Check proportional hazards assumption
cph.check_assumptions(final_cox_data, show_plots=True)
```

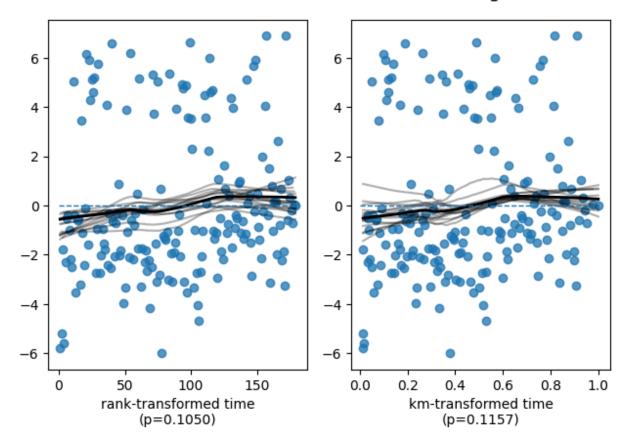
```
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
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Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
Bootstrapping lowess lines. May take a moment...
```

Bootstrapping lowess lines. May take a moment...

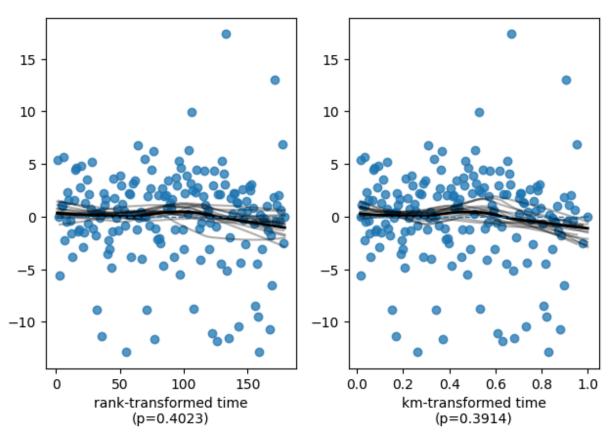
Proportional hazard assumption looks okay.

```
Out[32]: [[<Axes: xlabel='rank-transformed time\n(p=0.1050)'>,
            <Axes: xlabel='km-transformed time\n(p=0.1157)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.4023)'>,
            <Axes: xlabel='km-transformed time\n(p=0.3914)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.5798)'>,
            <Axes: xlabel='km-transformed time\n(p=0.6410)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.6326)'>,
            <Axes: xlabel='km-transformed time\n(p=0.6293)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.3914)'>,
           <Axes: xlabel='km-transformed time\n(p=0.4004)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.1004)'>,
            <Axes: xlabel='km-transformed time\n(p=0.1057)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.3749)'>,
            <Axes: xlabel='km-transformed time\n(p=0.3740)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.8431)'>,
            <Axes: xlabel='km-transformed time\n(p=0.8348)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.1995)'>,
            <Axes: xlabel='km-transformed time\n(p=0.2228)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.7497)'>,
            <Axes: xlabel='km-transformed time\n(p=0.7584)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.0920)'>,
            <Axes: xlabel='km-transformed time\n(p=0.1069)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.4158)'>,
            <Axes: xlabel='km-transformed time\n(p=0.4601)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.6103)'>,
            <Axes: xlabel='km-transformed time\n(p=0.6213)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.3416)'>,
            <Axes: xlabel='km-transformed time\n(p=0.3359)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.0410)'>,
            <Axes: xlabel='km-transformed time\n(p=0.0451)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.0181)'>,
            <Axes: xlabel='km-transformed time\n(p=0.0267)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.9159)'>,
            <Axes: xlabel='km-transformed time\n(p=0.7953)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.3657)'>,
            <Axes: xlabel='km-transformed time\n(p=0.3343)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.3457)'>,
            <Axes: xlabel='km-transformed time\n(p=0.3371)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.6208)'>,
            <Axes: xlabel='km-transformed time\n(p=0.6098)'>],
           [<Axes: xlabel='rank-transformed time\n(p=0.0274)'>,
            <Axes: xlabel='km-transformed time\n(p=0.0322)'>]]
```

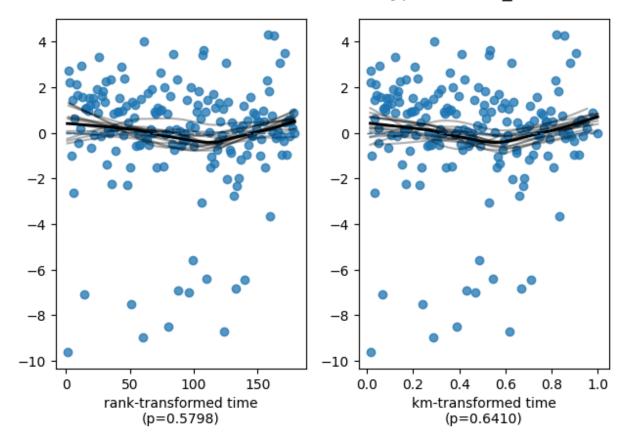
Scaled Schoenfeld residuals of 'Neuro signs'



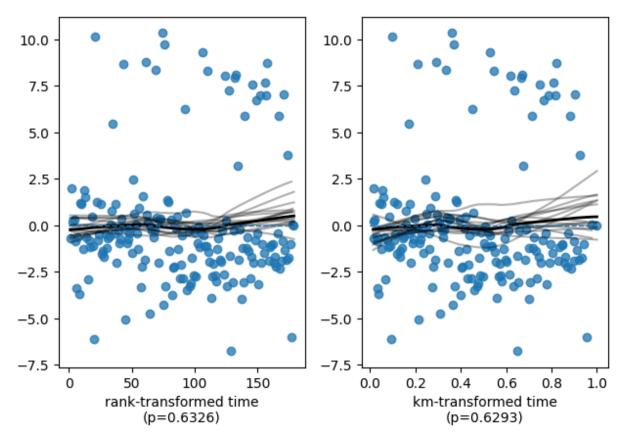
Scaled Schoenfeld residuals of 'Complications'



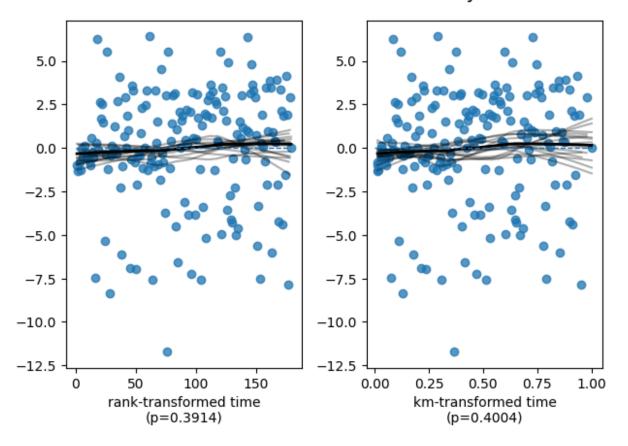
Scaled Schoenfeld residuals of 'Hypertensive_Yes4'



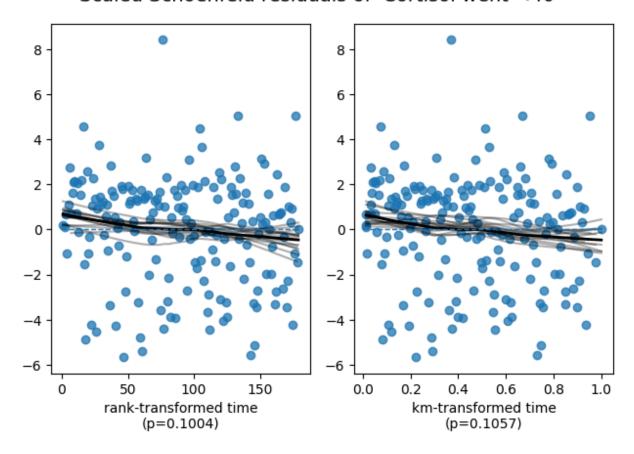
Scaled Schoenfeld residuals of 'Oversuppresion?'



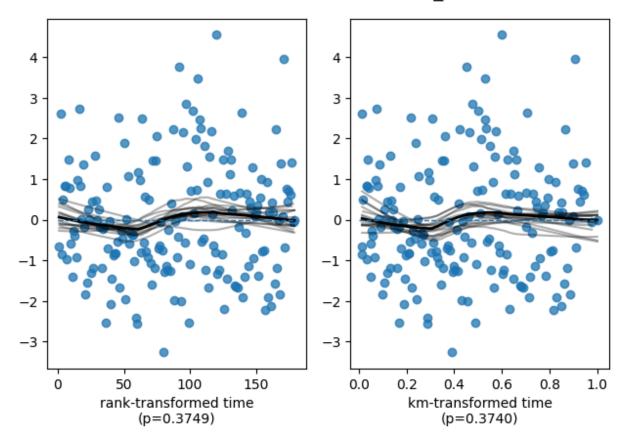
Scaled Schoenfeld residuals of 'Cortisol stayed <250'



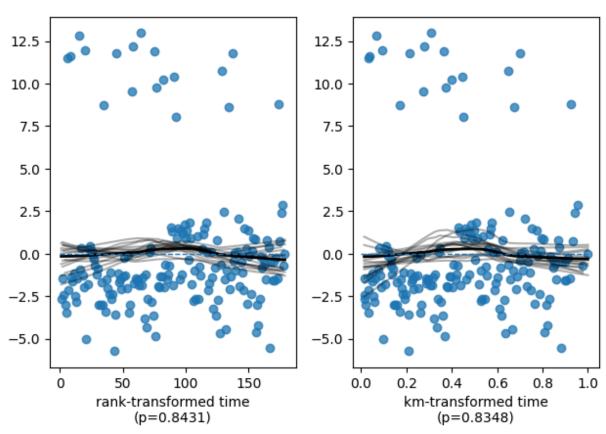
Scaled Schoenfeld residuals of 'Cortisol went <40'



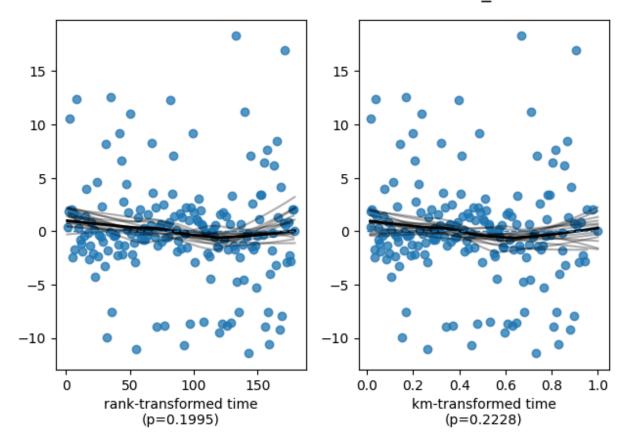
Scaled Schoenfeld residuals of 'Number_comorbidities'



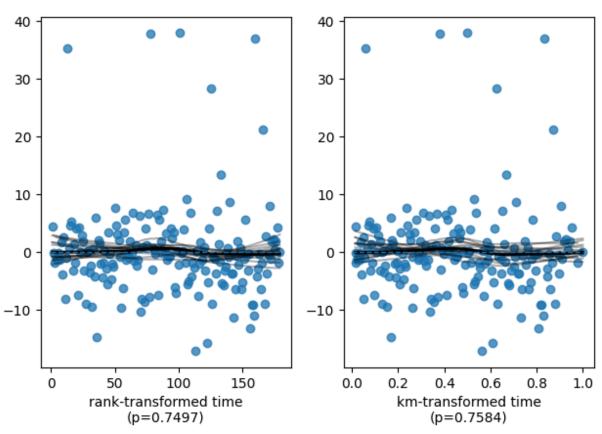
Scaled Schoenfeld residuals of 'comorb_UTI'



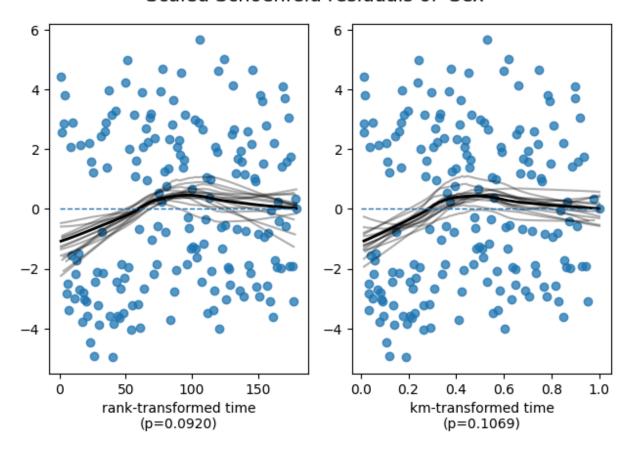
Scaled Schoenfeld residuals of 'comorb_dm'



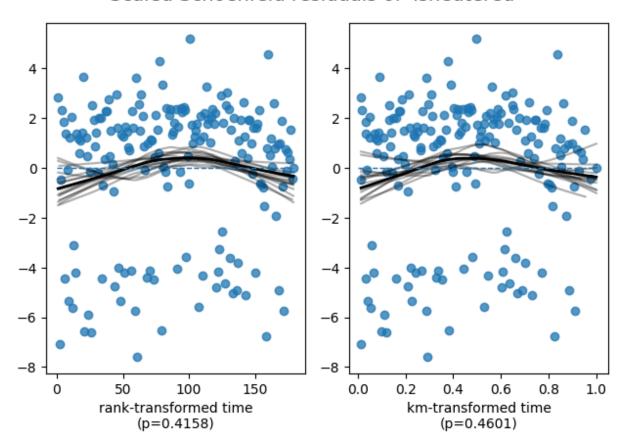
Scaled Schoenfeld residuals of 'comorb_hypot'



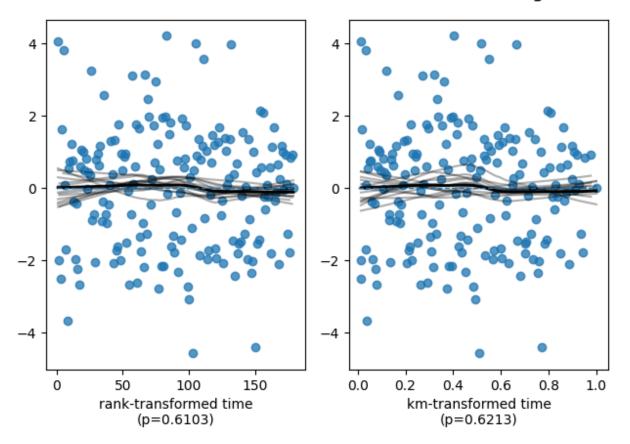
Scaled Schoenfeld residuals of 'Sex'



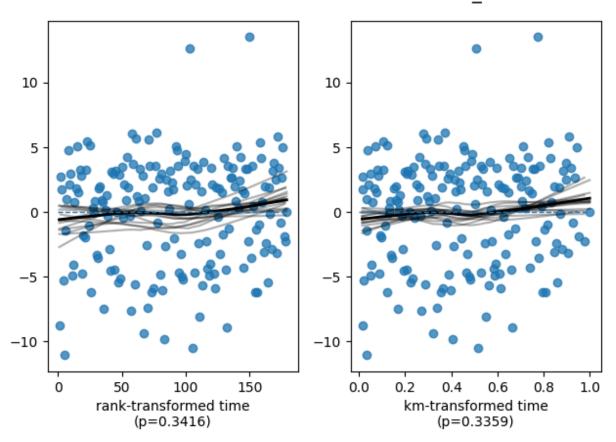
Scaled Schoenfeld residuals of 'Isneutered'



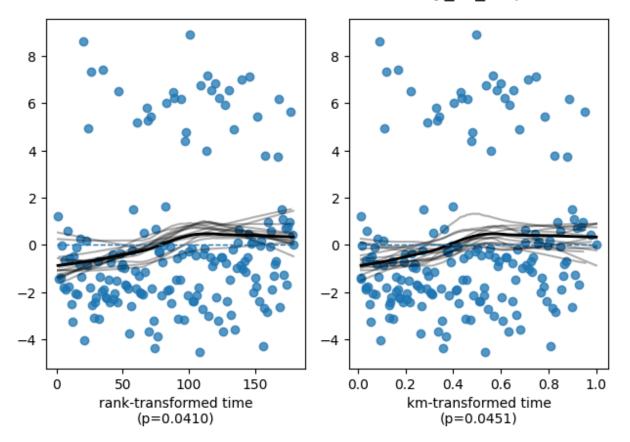
Scaled Schoenfeld residuals of 'BreedRelativeWeight'



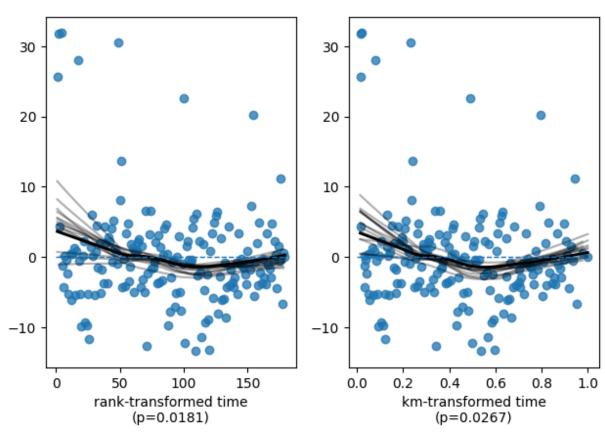
Scaled Schoenfeld residuals of 'Purebreed_status'



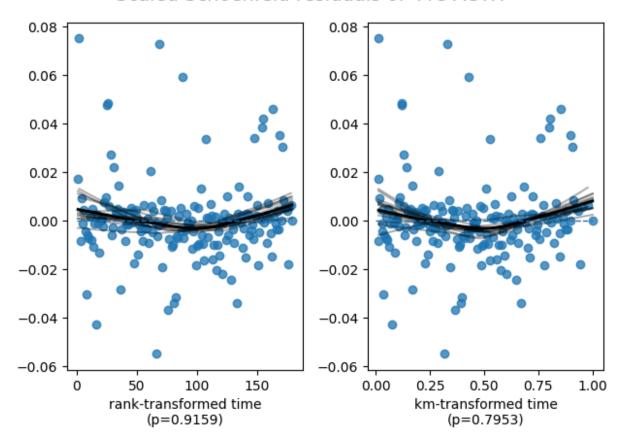
Scaled Schoenfeld residuals of 'Stay_vs_stop'



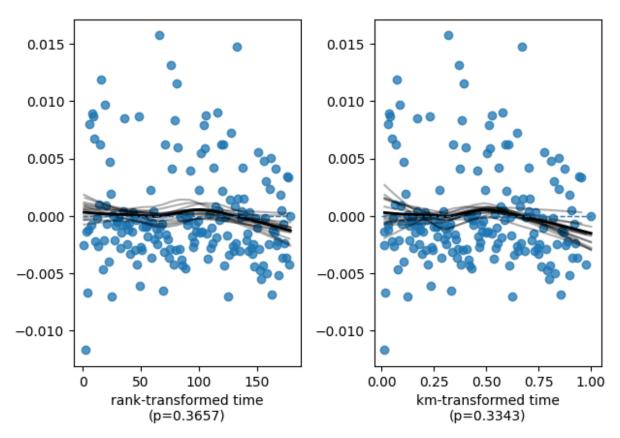
Scaled Schoenfeld residuals of 'Treated with trilostane_7'



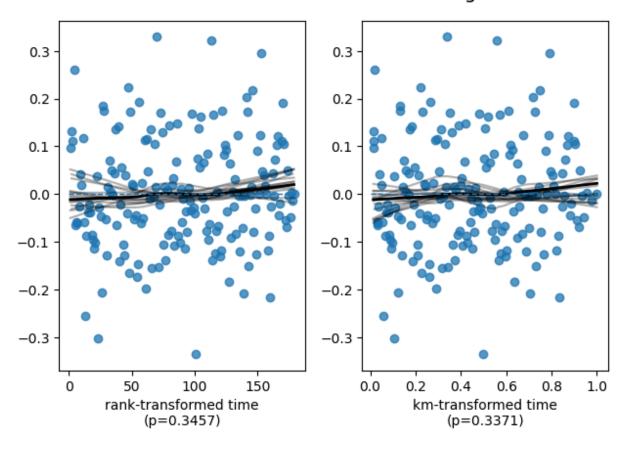
Scaled Schoenfeld residuals of 'Pre-ACTH'



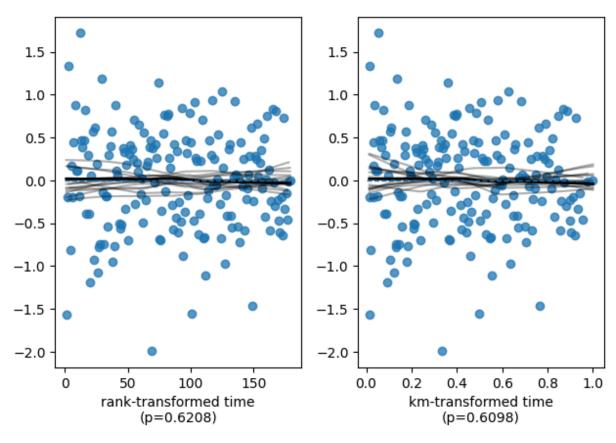
Scaled Schoenfeld residuals of 'Post-ACTH'



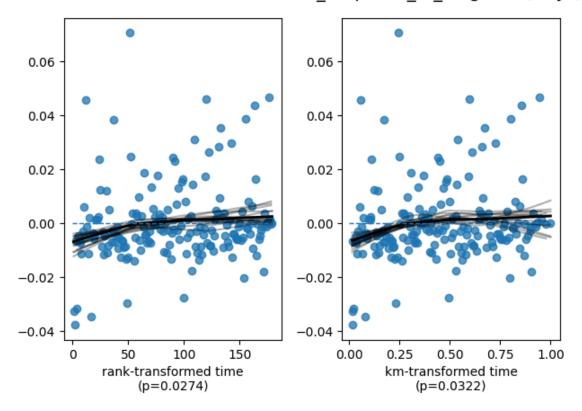
Scaled Schoenfeld residuals of 'weight'



Scaled Schoenfeld residuals of 'Age_at_diagnosis(Years)'

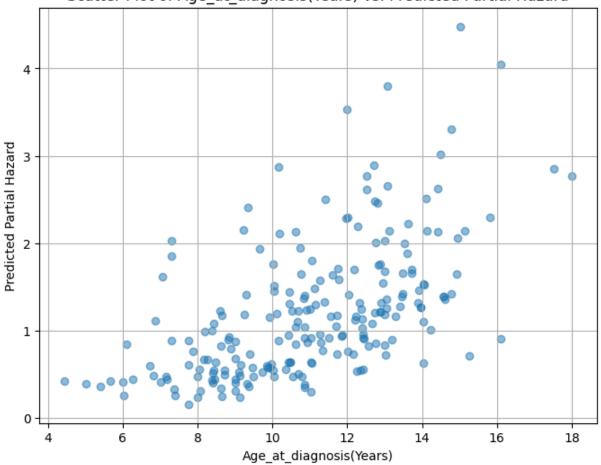


Scaled Schoenfeld residuals of 'Time suspicion to diagnosis(Days)'

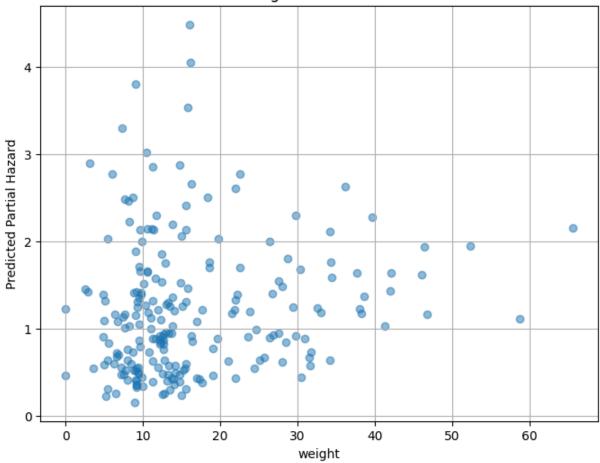


Proportional Hazards Assumption is Satisfied: The absence of any systematic trend in the residuals implies that the proportional hazards assumption is met. This is good news as it validates that the model is correctly specified in terms of the proportional hazards assumption.

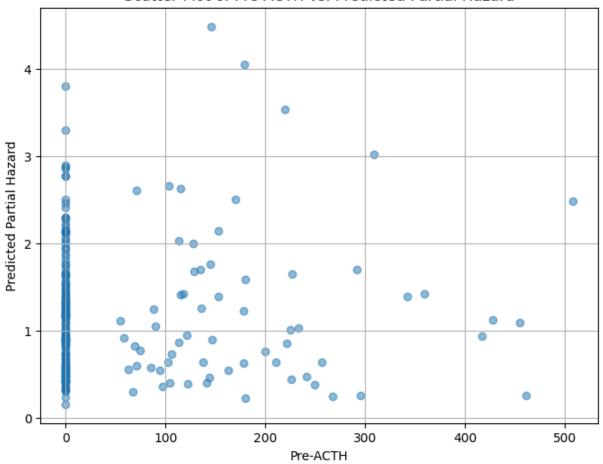
Scatter Plot of Age_at_diagnosis(Years) vs. Predicted Partial Hazard

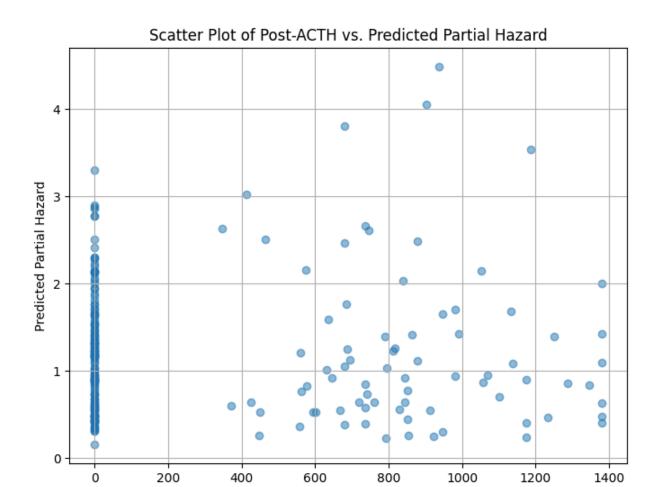






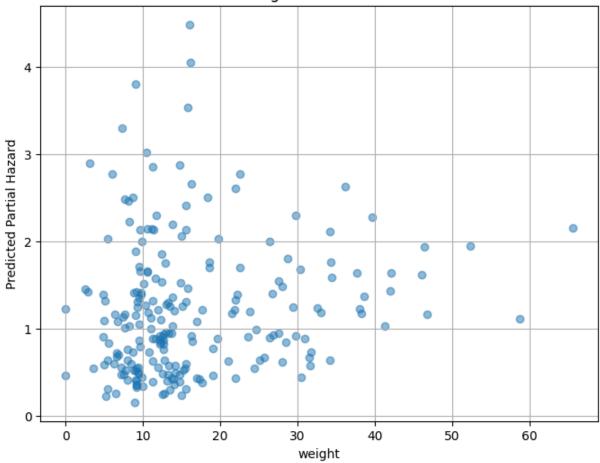
Scatter Plot of Pre-ACTH vs. Predicted Partial Hazard



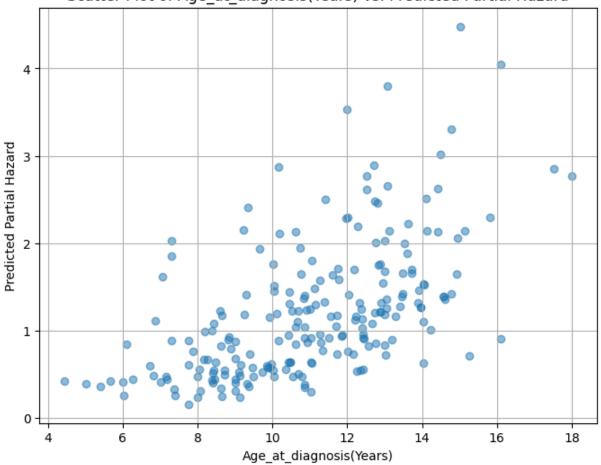


Post-ACTH

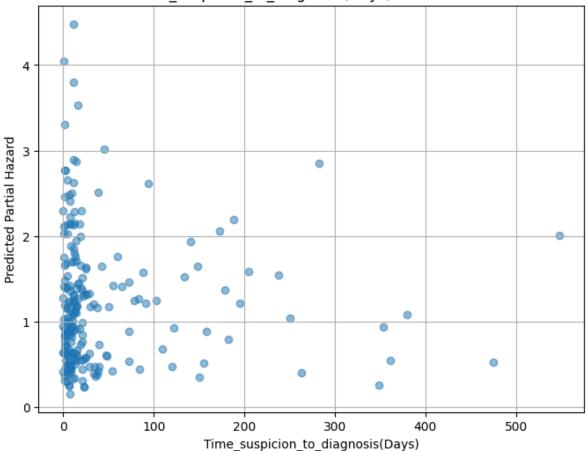




Scatter Plot of Age_at_diagnosis(Years) vs. Predicted Partial Hazard



Scatter Plot of Time suspicion to diagnosis(Days) vs. Predicted Partial Hazard



CoxPH for Comorbities

Subset: canine_comorb is used for the Analysis

```
# Creating a new DataFrame for comorbidities
In [34]:
         canine_comorb = pd.DataFrame()
         # Converting the categorical variables to binary
         canine_comorb['comorb_UTI'] = canine_subset['comorb_UTI'].map({'Yes': 1, 'No': 0})
         canine_comorb['comorb_dm'] = canine_subset['comorb_dm'].map({'Yes': 1, 'No': 0})
         canine_comorb['comorb_hypot'] = canine_subset['comorb_hypot'].map({'Yes': 1, 'No':
         canine_comorb['Number_comorbidities'] = canine_subset['Number_comorbidities']
         canine_comorb['Days_Diagnosis_to_Failure'] = canine_subset['Days_Diagnosis_to_Failu
         canine_comorb['Died'] = canine_subset['Died']
         # Displaying the modified DataFrame
         print(canine_comorb[['comorb_UTI','comorb_dm','comorb_hypot']])
         # Displaying the modified DataFrame
         print(canine_subset[['comorb_UTI','comorb_dm','comorb_hypot']])
         print("-----")
         print(canine_comorb.head())
```

```
comorb_UTI comorb_dm comorb_hypot
       0
                    0
       1
                             0
                                          0
       2
                   0
                            1
                                          0
       3
                    0
                             0
                                          0
       4
                   1
                              0
                                          0
                  . . .
       214
                   0
                              0
       215
                   0
                              0
       216
                   0
                             0
                                         0
       217
                    0
                              0
                                          0
                    0
       218
       [219 rows x 3 columns]
           comorb_UTI comorb_dm comorb_hypot
                   No
                            No
       0
                   No
                            No
       1
                                         No
       2
                  No
                           Yes
                                        No
       3
                  No
                           No
                                        No
       4
                  Yes
                           No
                                         No
                  . . .
       214
                  No
                           No
                                         No
       215
                  No
                           No
                                         No
       216
                 No
                           No
                                        No
       217
                   No
                           No
                                         No
       218
                   No
                           No
                                         No
       [219 rows x 3 columns]
         comorb_UTI comorb_dm comorb_hypot Number_comorbidities \
                                                           0.0
       1
                  0
                           0
                                        0
                                                           1.0
       2
                  0
                          1
                                      0
                                                           3.0
       3
                  0
                            0
                                        0
                                                           0.0
       4
                  1
                            0
                                                           1.0
          Days_Diagnosis_to_Failure Died
       0
                               1778
                               1842
       1
                                      0
       2
                               1260
                                      1
       3
                               942
                                      1
       4
In [35]: # Checking the data types
         print("Data types before fitting:")
         print(canine_comorb.dtypes)
         # Convert all categorical columns to numeric
         for col in canine_comorb.select_dtypes(include=['category']).columns:
             canine_comorb[col] = canine_comorb[col].cat.codes # Convert categorical to num
         # Checking the data types
         print("Data types after fitting:")
         print(canine_comorb.dtypes)
```

```
comorb_UTI
                                     category
        comorb_dm
                                     category
        comorb_hypot
                                     category
        Number_comorbidities
                                     category
        Days_Diagnosis_to_Failure
                                       int64
                                     category
        dtype: object
        Data types after fitting:
        comorb_UTI
                                      int8
        comorb_dm
                                      int8
        comorb_hypot
                                     int8
        Number_comorbidities
                                     int8
        Days_Diagnosis_to_Failure
                                     int64
        Died
                                      int8
        dtype: object
In [36]: # Fit multivariate model including all comorbidities and number of comorbidities
         cph.fit(canine_comorb[['Days_Diagnosis_to_Failure', 'Died',
                                 'comorb_UTI', 'comorb_dm',
                                 'comorb_hypot', 'Number_comorbidities']],
                 duration_col='Days_Diagnosis_to_Failure', event_col='Died')
         print("Multivariate analysis including all comorbidities:")
         cph.print_summary()
```

Multivariate analysis including all comorbidities:

Data types before fitting:

| model | lifelines.CoxPHFitter | | |
|---------------------------|-----------------------------|--|--|
| duration col | 'Days_Diagnosis_to_Failure' | | |
| event col | 'Died' | | |
| baseline estimation | breslow | | |
| number of observations | 219 | | |
| number of events observed | 179 | | |
| partial log-likelihood | -790.24 | | |
| time fit was run | 2024-11-23 18:26:10 UTC | | |

| | coef | exp(coef) | se(coef) | coef lower 95% | coef upper 95% | exp(coef) lower 95% | exp(coef) upper 95% | cmp to |
|----------------------|-------|-----------|----------|----------------------|----------------------|---------------------------|---------------------------|-----------|
| comorb_UTI | 0.60 | 1.82 | 0.25 | 0.10 | 1.09 | 1.11 | 2.97 | 0.00 |
| comorb_dm | 0.02 | 1.02 | 0.25 | -0.47 | 0.51 | 0.62 | 1.66 | 0.00 |
| comorb_hypot | 0.10 | 1.11 | 0.42 | -0.73 | 0.93 | 0.48 | 2.53 | 0.00 |
| Number_comorbidities | -0.10 | 0.90 | 0.08 | -0.26 | 0.05 | 0.77 | 1.06 | 0.00 |

Concordance 0.57
Partial AIC 1588.49
log-likelihood ratio test 6.62 on 4 df
-log2(p) of II-ratio test 2.67

In [37]: mpl.rcParams['figure.max_open_warning'] = 50
Check proportional hazards assumption
cph.check_assumptions(canine_comorb, show_plots=True)

Bootstrapping lowess lines. May take a moment...

Bootstrapping lowess lines. May take a moment...

Bootstrapping lowess lines. May take a moment...

Bootstrapping lowess lines. May take a moment...

The ``p_value_threshold`` is set at 0.01. Even under the null hypothesis of no viola tions, some

covariates will be below the threshold by chance. This is compounded when there are many covariates.

Similarly, when there are lots of observations, even minor deviances from the propor tional hazard

assumption will be flagged.

With that in mind, it's best to use a combination of statistical tests and visual tests to determine

the most serious violations. Produce visual plots using ``check_assumptions(..., sho w_plots=True)``

and looking for non-constant lines. See link [A] below for a full example.

| chi squared | null_distribution |
|--|--------------------|
| 1 | degrees_of_freedom |
| | model |
| proportional_hazard_test | test_name |

| | | test_statistic | р | -log2(p) |
|----------------------|------|----------------|--------|----------|
| Number_comorbidities | km | 9.92 | <0.005 | 9.26 |
| | rank | 10.21 | <0.005 | 9.48 |
| comorb_UTI | km | 0.05 | 0.82 | 0.28 |
| | rank | 0.05 | 0.83 | 0.27 |
| comorb_dm | km | 0.43 | 0.51 | 0.97 |
| | rank | 0.57 | 0.45 | 1.15 |
| comorb_hypot | km | 0.06 | 0.81 | 0.31 |
| | rank | 0.04 | 0.84 | 0.26 |

Variable 'Number_comorbidities' failed the non-proportional test: p-value is 0.00
 14.

Advice 1: the functional form of the variable 'Number_comorbidities' might be inc orrect. That is,

there may be non-linear terms missing. The proportional hazard test used is very sen sitive to

incorrect functional forms. See documentation in link [D] below on how to specify a functional form.

Advice 2: try binning the variable 'Number_comorbidities' using pd.cut, and then specify it in

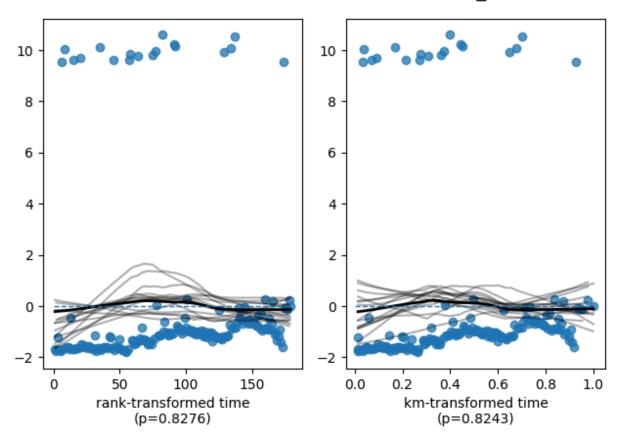
`strata=['Number_comorbidities', \dots]` in the call in `.fit`. See documentation in l ink [B] below.

Advice 3: try adding an interaction term with your time variable. See documentati on in link [C] below.

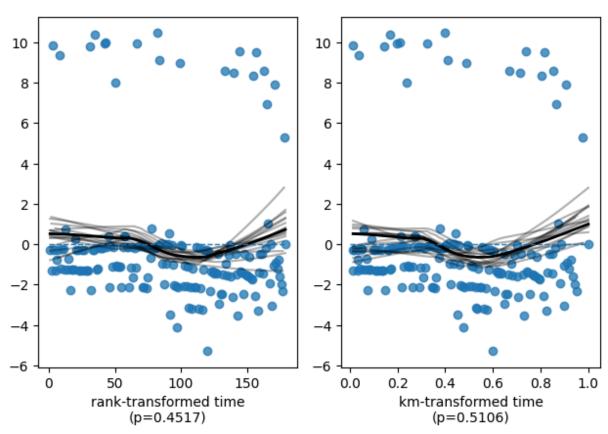
- -

- [A] https://lifelines.readthedocs.io/en/latest/jupyter_notebooks/Proportional%20haz ard%20assumption.html
- [B] https://lifelines.readthedocs.io/en/latest/jupyter_notebooks/Proportional%20haz ard%20assumption.html#Bin-variable-and-stratify-on-it
- [C] https://lifelines.readthedocs.io/en/latest/jupyter_notebooks/Proportional%20haz ard%20assumption.html#Introduce-time-varying-covariates
- [D] https://lifelines.readthedocs.io/en/latest/jupyter_notebooks/Proportional%20haz ard%20assumption.html#Modify-the-functional-form
- [E] https://lifelines.readthedocs.io/en/latest/jupyter_notebooks/Proportional%20haz ard%20assumption.html#Stratification

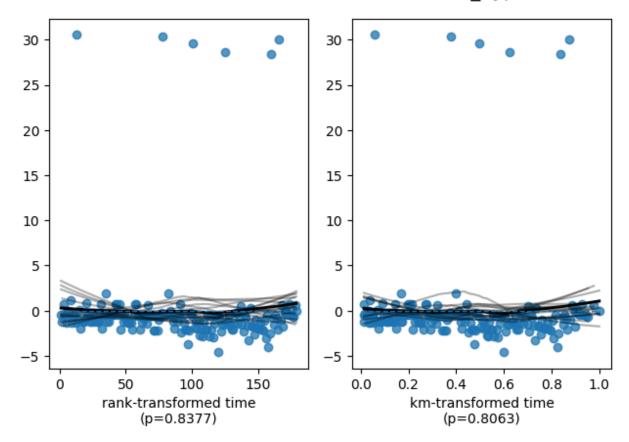
Scaled Schoenfeld residuals of 'comorb_UTI'



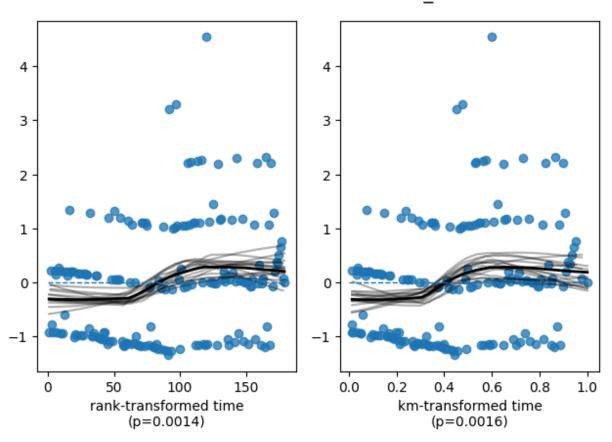
Scaled Schoenfeld residuals of 'comorb_dm'



Scaled Schoenfeld residuals of 'comorb_hypot'



Scaled Schoenfeld residuals of 'Number_comorbidities'



Random Survival Forests

Subset: random_data is used for the analysis

```
In [38]:
        # Creating random_data df for the Random Survival Forest Model
         random_data = cox_data[['Days_Diagnosis_to_Failure',
                                   'Died', 'Neuro signs', 'Complications',
                                   'Hypertensive_Yes4', 'Oversuppresion?',
                                   'Cortisol stayed <250', 'Cortisol went <40',
                                   'Number_comorbidities', 'comorb_UTI',
                                   'comorb_dm', 'comorb_hypot', 'Sex',
                                   'Isneutered', 'BreedRelativeWeight',
                                   'Purebreed_status', 'Stay_vs_stop',
                                   'Treated with trilostane_7', 'Pre-ACTH',
                                   'Post-ACTH', 'weight',
                                   'Age_at_diagnosis(Years)',
                                   'Time_suspicion_to_diagnosis(Days)']]
         # Checking the first few rows to ensure it's set up correctly
         print(random_data.head())
```

```
Days_Diagnosis_to_Failure Died Neuro signs Complications \
                                                    0
        0
                                1778
                                                    0
                                                                   2
        1
                                1842
                                         0
        2
                                                                   2
                                1260
                                         1
                                                    0
        3
                                 942
                                         1
                                                     0
                                                                   2
        4
                                 779
                                         1
                                                     0
                                                                   1
          Hypertensive_Yes4 Oversuppresion? Cortisol stayed <250 Cortisol went <40 \
                          2
                                          1
                                                               1
                          2
                                          1
                                                               2
                                                                                 2
        1
        2
                          2
                                          1
                                                               1
                                                                                 0
                          2
        3
                                          1
                                                               0
                                                                                 0
        4
           Number comorbidities comorb UTI ... Isneutered BreedRelativeWeight
        0
                              0
                                         0 ...
                                                         1
                              1
                                                                             1
        1
                                           . . .
                                                         1
                                                                             2
        2
                              3
                                         0 ...
                                                         1
        3
                              0
                                                         1
                                                                             1
                                           . . .
        4
                              1
                                                                             2
           Purebreed_status Stay_vs_stop Treated with trilostane_7 Pre-ACTH \
        0
                          1
                                       1
                                                                  1
                                                                          0.0
                                                                          0.0
        1
                          1
                                       1
                                                                  1
        2
                          1
                                       1
                                                                  1
                                                                        267.0
        3
                          1
                                       2
                                                                  1
                                                                          0.0
                                       2
                                                                  1
                                                                        145.0
        4
                          1
           Post-ACTH weight Age_at_diagnosis(Years) \
        0
                 0.0
                      7.55
                                                10.04
                      7.20
        1
                 0.0
                                                10.36
                     12.50
        2
               922.0
                                                8.66
                 0.0 19.00
        3
                                                10.79
               684.0 34.25
                                                10.03
        4
           Time_suspicion_to_diagnosis(Days)
        0
                                          22
        1
        2
                                           6
        3
                                          35
        4
                                          60
        [5 rows x 23 columns]
In [39]: # Preparing the features (X) and target (y)
         X = random_data.drop(columns=['Days_Diagnosis_to_Failure', 'Died'])
         # Creating the target variable with duration and event indicator
         y = pd.DataFrame({'event': random_data['Died'].astype(bool), 'time': random_data['
         y = y.to_records(index=False)
         # Creating and fitting the Random Survival Forest model
         rsf = RandomSurvivalForest(n_estimators=1000, min_samples_split=10, min_samples_lea
         rsf.fit(X, y)
```

```
Out[39]:

RandomSurvivalForest

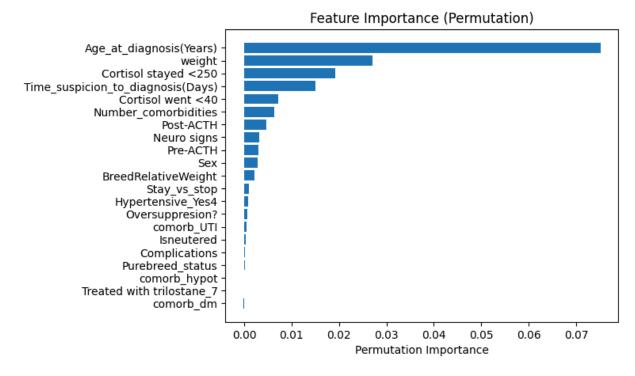
RandomSurvivalForest(min_samples_leaf=15, min_samples_split=10, n_estimators=1000, random_state=55)
```

```
In [40]: # Calculating the concordance index
c_index = concordance_index_censored(y['event'], y['time'], rsf.predict(X))
print(f"Concordance Index: {c_index[0]}")
```

Concordance Index: 0.7055767054223686

```
In [41]: # Calculating the permutation importance
    result = permutation_importance(rsf, X, y, n_repeats=10, random_state=55)

# Displaying the feature importance from permutation importance
    sorted_idx = result.importances_mean.argsort()
    plt.barh(range(len(sorted_idx)), result.importances_mean[sorted_idx], align='center
    plt.yticks(range(len(sorted_idx)), X.columns[sorted_idx])
    plt.xlabel("Permutation Importance")
    plt.title("Feature Importance (Permutation)")
    plt.show()
```



```
In [42]: # Calculating the permutation importance
# Feature Importance not available in this package
result = permutation_importance(rsf, X, y, n_repeats=10, random_state=55)

# Creating a DataFrame to display the results
importance_df = pd.DataFrame({
    'Feature': X.columns,
    'Importance Mean': result.importances_mean,
    'Importance Std': result.importances_std
})
```

```
# Sorting by importance
importance_df = importance_df.sort_values(by='Importance Mean', ascending=False)
# Displaying the feature importance
print(importance_df)
```

```
Feature Importance Mean Importance Std
19
              Age_at_diagnosis(Years)
                                               0.075152
                                                               0.018984
18
                               weight
                                               0.027076
                                                               0.008896
4
                 Cortisol stayed <250
                                               0.019158
                                                               0.011958
20 Time_suspicion_to_diagnosis(Days)
                                                               0.004379
                                               0.015084
5
                    Cortisol went <40
                                               0.007233
                                                               0.009773
6
                 Number_comorbidities
                                               0.006354
                                                               0.002053
17
                            Post-ACTH
                                               0.004651
                                                               0.001002
0
                          Neuro signs
                                               0.003097
                                                               0.002291
16
                             Pre-ACTH
                                               0.002974
                                                               0.000757
10
                                  Sex
                                               0.002799
                                                               0.000702
12
                  BreedRelativeWeight
                                               0.002120
                                                               0.000745
14
                         Stay_vs_stop
                                               0.001065
                                                               0.000501
2
                    Hypertensive_Yes4
                                               0.000859
                                                               0.000336
3
                      Oversuppresion?
                                               0.000720
                                                               0.001124
7
                           comorb_UTI
                                               0.000540
                                                               0.000252
                           Isneutered
                                               0.000417
                                                               0.000500
11
                                               0.000180
                                                               0.000766
1
                        Complications
13
                     Purebreed_status
                                               0.000129
                                                               0.000457
9
                         comorb hypot
                                               0.000000
                                                               0.000000
15
            Treated with trilostane_7
                                               0.000000
                                                               0.000000
8
                            comorb_dm
                                              -0.000159
                                                               0.000071
```

```
In [43]: # Predicting the survival function for each individual in X
    predicted_survival = rsf.predict_survival_function(X)

# Visualization of survival functions for each individual
    for i in range(len(predicted_survival)):
        plt.step(predicted_survival[i].x, predicted_survival[i].y, where="post", label="plt.title("Random Survival Forest - Survival Function")
    plt.xlabel("Time")
    plt.ylabel("Survival Probability")
    plt.show()
```

Random Survival Forest - Survival Function

