

## ***In silico* analysis of the PTCome distribution (Kernel plots)**

### **1. Potential PTCome**

- a. To obtain the potential PTCome, extract from Gene database (NCBI, <https://www.ncbi.nlm.nih.gov/gene/>) the cDNA sequence of the desired gene. Copy it in a text document.
- b. Import the text document in the PTCMAKER program or run the online code.
- c. Run the code in (<https://github.com/leiretorices/Kernel-plots>, section 1) to obtain unique potential PTC residues.

### **2. Germline-associated PTCome**

- a. To obtain the germline-associated PTCome, browse for the desired PTP in the HGMD database (<https://www.hgmd.cf.ac.uk>). HGMD displays a data frame with missense and nonsense mutations together.
- b. Import the downloaded file in RStudio by File > Import Dataset > From Excel, or using the `read_excel()` function.
- c. Run the code in (<https://github.com/leiretorices/Kernel-plots>, section 2) to obtain unique PTC mutated residues.

### **3. Kernel plot representation**

- a. Insert the amino acid length and the name of your protein of interest.
- b. Run the code in (<https://github.com/leiretorices/Kernel-plots>, section 3) to obtain a kernel plot representation. The three vectors (potential PTCome, cancer-associated PTCome and germline-associated PTCome) are rescaled to 100 to facilitate comparisons between proteins. Mirror vectors are generated to avoid bias at boundaries and a density is calculated of sum of the parental vector and the mirror vectors. The final plot contains black, and blue curves, which represent the potential PTCome and the germline-associated PTCome, respectively. Cancer-associated PTCome (COSMIC database) can be obtained from <https://github.com/leiretorices/Kernel-plots> section 4.