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Independence tests between two point processes. Application to the study of neuronal spike trains synchronization

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Introduction of the biological context

Transmission of the neuronal message

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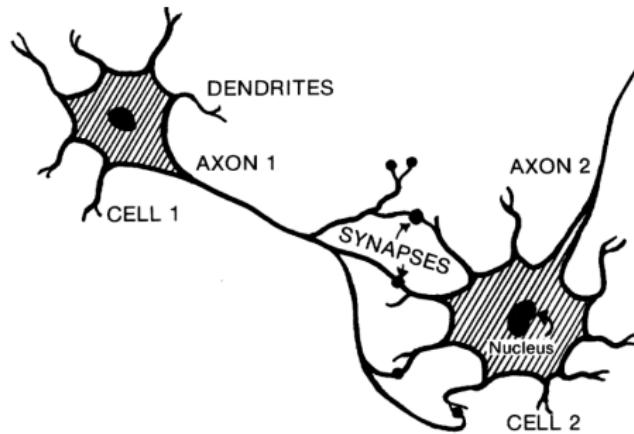
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The neuronal information is transmitted by the spikes/action potentials.



Electric potential of the cell

Synaptic Integration

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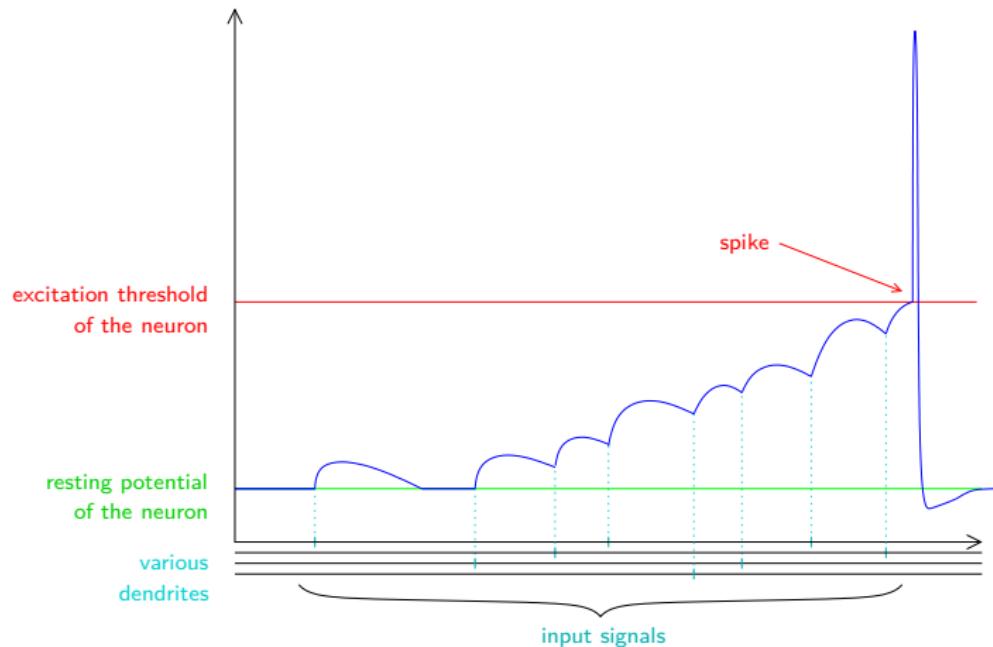
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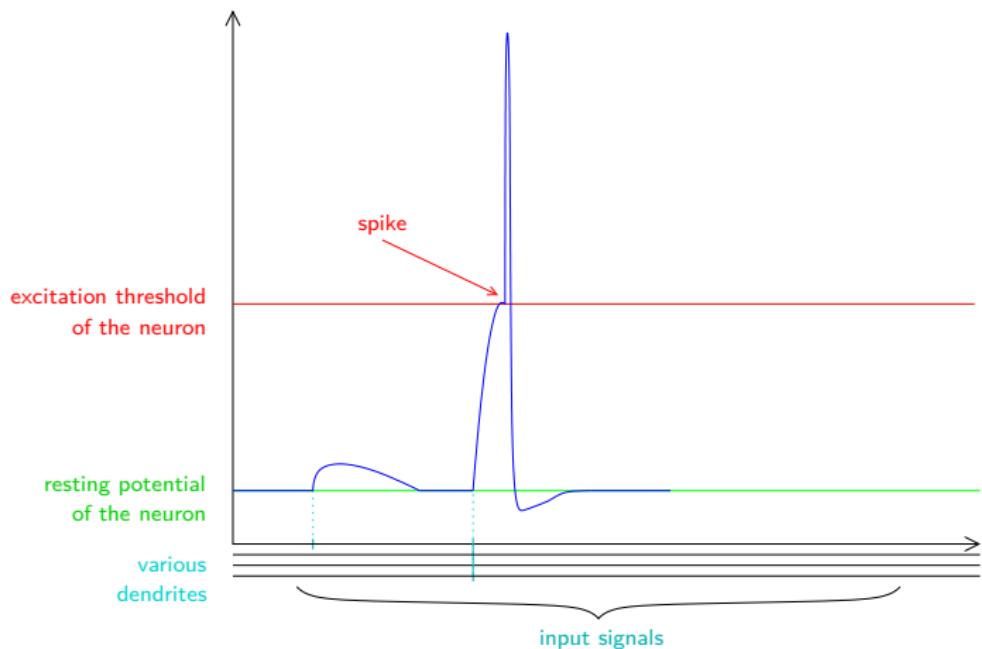
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[Grammont and Riehle (1999)]: neurons coordinate their activity at very precise moments.

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Problematic

Detection of dependences between neurons.

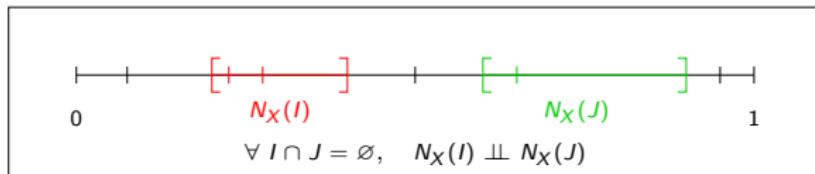
Statistical Modeling for Neuronal Activity

Spike = "all-or-none" phenomenon,
⇒ Model: point processes on a time slot $[0, 1]$.

Definition

Point process on $[0, 1]$ = random countable set of points in $[0, 1]$.
 \mathcal{X} := the set of almost surely finite Point process on $[0, 1]$.

Example: X homogeneous Poisson process with intensity $\lambda > 0$.



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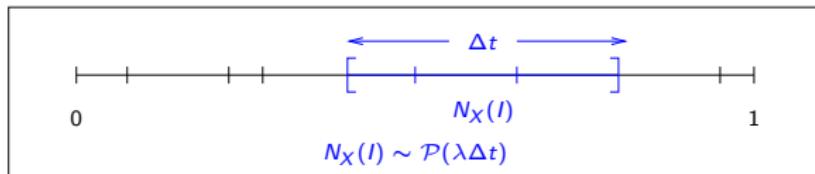
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$\mathcal{X} :=$ the set of almost surely finite Point process on $[0, 1]$.

For X a point process in $[0, 1]$, we introduce

$$dN_X = \sum_{T \in X} \delta_T.$$

Notion of coincidence

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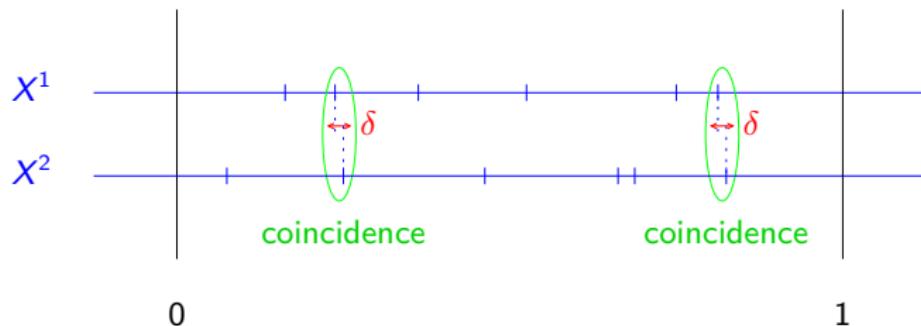
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Notion of (delayed) coincidence [Grün et al. (1999)]

$\varphi_{\delta}^{\text{coinc}}$ counts the number of coincidences between two point processes:

$$\varphi_{\delta}^{\text{coinc}}(X^1, X^2) = \int_a^b \int_a^b \mathbb{1}_{\{|u-v| \leq \delta\}} dN_{X^1}(u) dN_{X^2}(v).$$



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Unitary Events [Grün et al. (1999)]

Most well known method using delayed coincidences.

Problem:

- Very few methods theoretically justified.

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Unitary Events [Grün et al. (1999)]

Most well known method using delayed coincidences.

A correction from Tuleau-Malot et al. (2012)

Homogeneous Poisson process assumption \Rightarrow Determination of the asymptotic distribution of the test statistic.

Problems:

- Homogeneous Poisson process assumption questionable,
- Too few data to use the asymptotic distribution.

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Aim

To construct a non-asymptotic independence test between point processes with no assumption on the underlying distribution.

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Pipa et al. (2003)

Trial shuffling for another notion of coincidences.

Independence Test

Definition of the test statistic

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X^m : activity of neuron $m \in \{1, 2\}$ on $[0, 1]$.

$$(H_0) : X^1 \perp\!\!\!\perp X^2 \quad \text{against} \quad (H_1) : X^1 \not\perp\!\!\!\perp X^2$$

Often, in \mathbb{R}^k , tests based on $\iint \varphi(x^1, x^2) (dP_{(x^1, x^2)} - dP_{x^1}^1 dP_{x^2}^2)$.

For φ well chosen,

joint distribution \leftrightarrow product of marginals.

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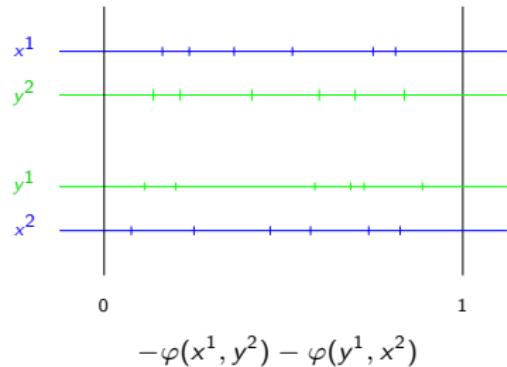
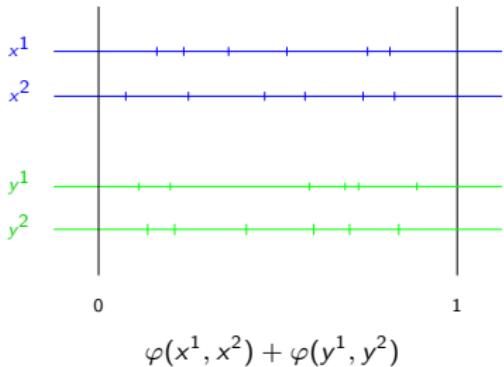
Often, in \mathbb{R}^k , tests based on $\iint \varphi(x^1, x^2) (dP_{(x^1, x^2)} - dP_{x^1}^1 dP_{x^2}^2)$.

Let $\mathbf{X}_n = (X_i)_{1 \leq i \leq n}$, where $X_i = (X_i^1, X_i^2)$ i.i.d. $\sim P$ in $\mathcal{X} \times \mathcal{X}$.

Unbiased estimator:

$$\frac{1}{n(n-1)} \sum_{i \neq i'} (\varphi(X_i^1, X_i^2) - \varphi(X_i^1, X_{i'}^2))$$

Let $h_\varphi((x^1, x^2), (y^1, y^2)) = \varphi(x^1, x^2) + \varphi(y^1, y^2) - \varphi(x^1, y^2) - \varphi(y^1, x^2)$,



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Let $h_\varphi((x^1, x^2), (y^1, y^2)) = \varphi(x^1, x^2) + \varphi(y^1, y^2) - \varphi(x^1, y^2) - \varphi(y^1, x^2)$,

the previous estimator becomes $\frac{1}{2n(n-1)} \sum_{i \neq j} h(X_i, X_j)$.

Test Statistic

$$\sqrt{n} U_{n,h}(\mathbb{X}_n) = \frac{\sqrt{n}}{n(n-1)} \sum_{i \neq j} h(X_i, X_j),$$

for $h : (\mathcal{X} \times \mathcal{X})^2 \rightarrow \mathbb{R}$ well chosen.

Bootstrap approach

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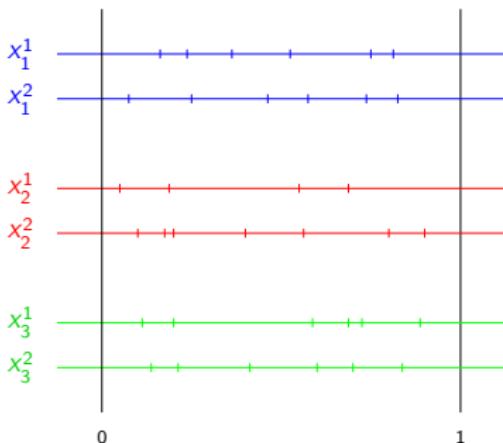
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Bootstrap approach of Romano (1988)

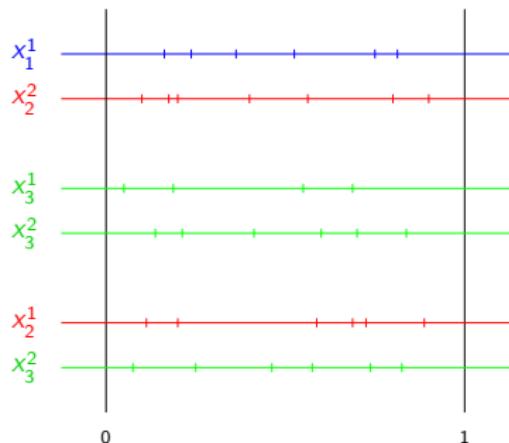
Given $\mathbb{X}_n = (X_i)_{1 \leq i \leq n}$, where $X_i = (X_i^1, X_i^2)$ i.i.d. $\sim P$ in $\mathcal{X} \times \mathcal{X}$.

The bootstrap sample is

$$\mathbb{X}_n^* = (X_{n,1}^*, \dots, X_{n,n}^*) \text{ i.i.d. } \sim P_n^1 \otimes P_n^2 = \left(\frac{1}{n} \sum_{1 \leq i \leq n} \delta_{X_i^1} \right) \otimes \left(\frac{1}{n} \sum_{1 \leq j \leq n} \delta_{X_j^2} \right).$$



$\sqrt{n}U_{n,h}(\mathbb{X}_n)$: on original data



$\sqrt{n}U_{n,h}(\mathbb{X}_n^*)$: on bootstrapped data

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Theorem 1

Let $h : (\mathcal{X} \times \mathcal{X})^2 \rightarrow \mathbb{R}$ such that $(\mathcal{A}_{Exp}^{Cent})$, $(\mathcal{A}_{Emp}^{Cent})$, $(\mathcal{A}_{brk}^{Mmt})$ and (\mathcal{A}^{Cont}) hold, then

$$d_{W_2} \left(\mathcal{L} \left(\sqrt{n} U_{n,h}; P_n^1 \otimes P_n^2 | \mathbb{X}_n \right), \mathcal{L} \left(\sqrt{n} U_{n,h}; P^1 \otimes P^2 \right) \right) \xrightarrow{n \rightarrow \infty} 0$$

P -a.s. in $(X_n)_n$.

Here, $\mathcal{L} \left(\sqrt{n} U_{n,h}; Q \right)$ denotes the distribution of $\sqrt{n} U_{n,h} (\mathbb{Z}_n)$ for $\mathbb{Z}_n = (Z_1, \dots, Z_n)$ sample of i.i.d. random variables with distribution Q .

Bootstrap test of independence

$$\Phi_{h,\alpha}^+ = \mathbb{1}_{\{\sqrt{n}U_{n,h}(\mathbb{X}_n) > q_{h,1-\alpha}^*(\mathbb{X}_n)\}}$$

where $q_{h,1-\alpha}^*(\mathbb{X}_n)$ is the $(1-\alpha)$ -quantile of $\mathcal{L}(\sqrt{n}U_{n,h}; P_n^1 \otimes P_n^2 | \mathbb{X}_n)$.

Theorem 2

- *Asymptotic level:*

Under (H_0) ,

$$\mathbb{P}(\Phi_{h,\alpha}^+ = 1) \xrightarrow[n \rightarrow \infty]{} \alpha$$

- *Asymptotic power:*

For any alternative P such that $\mathbb{E}[h(X_i, X_{i'})] > 0$,

$$\mathbb{P}(\Phi_{h,\alpha}^+ = 1) \xrightarrow[n \rightarrow +\infty]{} 1$$

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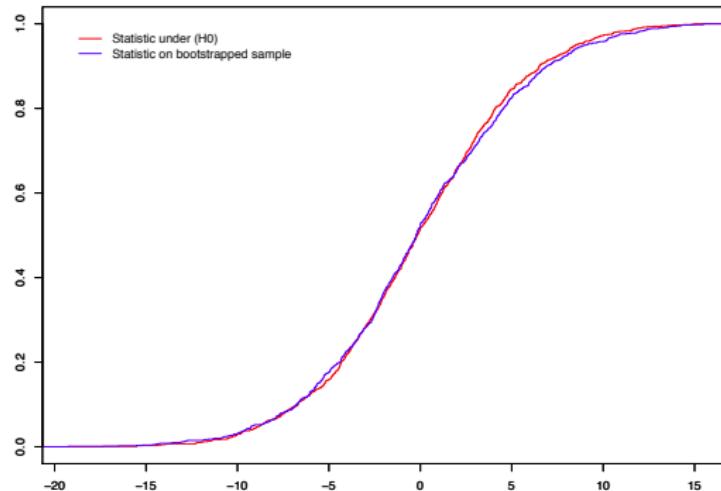
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Corollary of Theorem 1

Under (H_0) , $P = P^1 \otimes P^2$, and P -a.s. in $(X_n)_n$,

$$d_{W_2} \left(\mathcal{L} \left(\sqrt{n} U_{n,h}; P_n^1 \otimes P_n^2 | \mathbb{X}_n \right), \mathcal{L} \left(\sqrt{n} U_{n,h}; P \right) \right) \xrightarrow{n \rightarrow \infty} 0.$$



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Centered Gaussian limit of $\sqrt{n} U_{n,h}$ under independence

Under (H_0) ,

$$\mathcal{L} \left(\sqrt{n} U_{n,h}; P \right) \xrightarrow{n \rightarrow \infty} \mathcal{N} \left(0, \sigma_P^2 \right).$$

Quantiles

$q_{h,1-\alpha}^*(\mathbb{X}_n)$ converges a.s to the $(1-\alpha)$ -quantile of $\mathcal{N} \left(0, \sigma_P^2 \right)$.

Finally,

$$\mathbb{P} \left(\Phi_{h,\alpha}^+ = 1 \right) = \mathbb{P} \left(\sqrt{n} U_{n,h}(\mathbb{X}_n) > q_{h,1-\alpha}^*(\mathbb{X}_n) \right) \xrightarrow{n \rightarrow \infty} \alpha.$$

- LLN for U -statistics :
if $\mathbb{E} [|h(X, X')|] < +\infty$, then

$$U_{n,h}(\mathbb{X}_n) \xrightarrow[n \rightarrow \infty]{} \mathbb{E} [h(X, X')] > 0, \quad P\text{-a.s. in } (X_n)_n.$$

- Using the convergence of the quantile,

$$\frac{q_{h,1-\alpha}^*(\mathbb{X}_n)}{\sqrt{n}} \xrightarrow[n \rightarrow \infty]{} 0.$$

So finally,

$$\mathbb{P}(\Phi_{h,\alpha}^+ = 1) = \mathbb{P}\left(U_{n,h}(\mathbb{X}_n) > \frac{q_{h,1-\alpha}^*(\mathbb{X}_n)}{\sqrt{n}}\right) \xrightarrow[n \rightarrow +\infty]{} 1.$$

Basic Permutation Test

Idea of permutation

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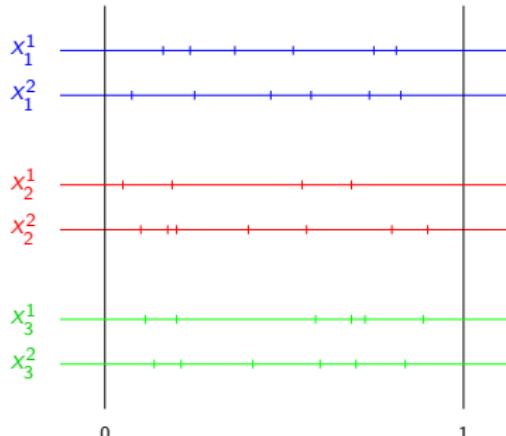
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Permuted sample

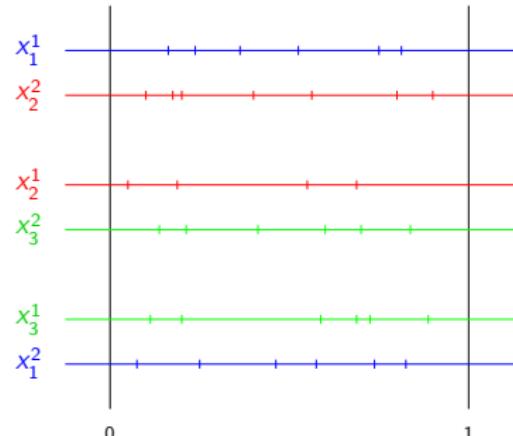
Given $\mathbb{X}_n = (X_i)_{1 \leq i \leq n}$, where $X_i = (X_i^1, X_i^2)$ i.i.d. $\sim P$ in $\mathcal{X} \times \mathcal{X}$.

The permuted sample is

$$\mathbb{X}_n^{s_n} = (X_1^{s_n}, \dots, X_n^{s_n}), \quad \text{with } X_i^{s_n} = (X_i^1, X_{s_n(i)}^2).$$



$\sqrt{n}U_{n,h}(\mathbb{X}_n)$: on original data



$\sqrt{n}U_{n,h}(\mathbb{X}_n^{s_n})$: on permuted data (with $s_n \sim \mathcal{U}(\mathfrak{S}_n)$ independent of \mathbb{X}_n)

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$$\mathbb{X}_n^{s_n} = (X_1^{s_n}, \dots, X_n^{s_n}), \quad \text{with } X_i^{s_n} = (X_i^1, X_{s_n(i)}^2).$$

Consistency

Let $s_n \sim \mathcal{U}(\{1, \dots, n\}) \perp\!\!\!\perp \mathbb{X}_n$.

Under (H_0) , $\mathbb{X}_n^{s_n}$, and \mathbb{X}_n have the same distribution.

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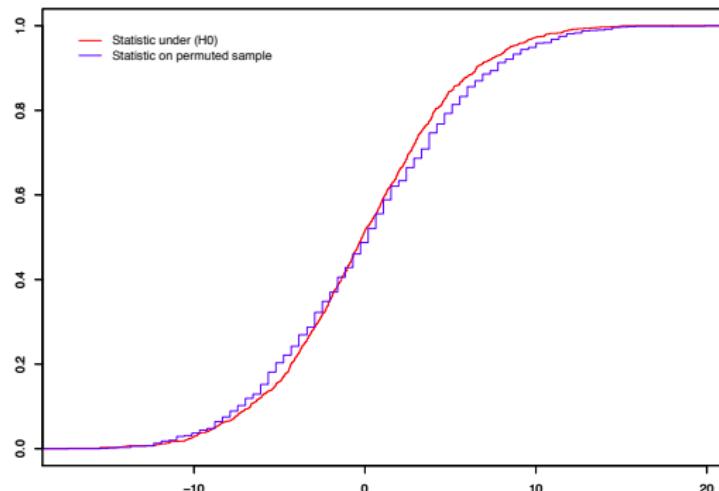
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Theorem 2

Let $h = h_\varphi$ with $\varphi : \mathcal{X}^2 \rightarrow \mathbb{R}$ satisfying $(\mathcal{A}^{Maj, \varphi})$ and $(\mathcal{A}_{Card}^{Mmt})$.
Then, under (H_0) ,

$$d_{W_2} \left(\mathcal{L} \left(\sqrt{n} U_{n, h_\varphi} (\mathbf{X}_n^s) | \mathbf{X}_n \right), \mathcal{L} \left(\sqrt{n} U_{n, h} ; P^1 \otimes P^2 \right) \right) \xrightarrow[n \rightarrow \infty]{\mathbb{P}} 0.$$



Permutation test of independence

$$\Psi_{h,\alpha}^+ = \mathbb{1}_{\left\{ \sqrt{n}U_{n,h_\varphi}(\mathbb{X}_n) > q_{h_\varphi,1-\alpha}^\pi(\mathbb{X}_n) \right\}}$$

where $q_{h_\varphi,1-\alpha}^\pi(\mathbb{X}_n)$ is the $(1-\alpha)$ -quantile of $\mathcal{L}\left(\sqrt{n}U_{n,h_\varphi}(\mathbb{X}_n^s) | \mathbb{X}_n\right)$.

Theorem 2

- *Asymptotic level:*
Under (H_0) ,

$$\mathbb{P}\left(\Psi_{h_\varphi,\alpha}^+ = 1\right) \leq \alpha,$$

and

$$\mathbb{P}\left(\Psi_{h_\varphi,\alpha}^+ = 1\right) \xrightarrow{n \rightarrow \infty} \alpha.$$

- *Asymptotic power:*
For any alternative P such that $\mathbb{E}[h(X_i, X_{i'})] > 0$,

$$\mathbb{P}\left(\Psi_{h_\varphi,\alpha}^+ = 1\right) \xrightarrow{n \rightarrow +\infty} 1.$$

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