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Online multiple testing with super-uniformity reward

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Abstract: Valid online inference is an important problem in contemporary multiple testing research, to which various solutions have been proposed recently. It is well-known that these existing methods can suffer from a significant loss of power if the null p-values are conservative. In this work, we extend the previously introduced methodology to obtain more powerful procedures for the case of super-uniformly distributed p-values. These types of p-values arise in important settings, e.g. when discrete hypothesis tests are performed. To this end, we introduce the method of super-uniformity reward (SUR) that incorporates information about the individual null cumulative distribution functions. Our approach yields several new 'rewarded' procedures that offer uniform power improvements over known procedures and come with mathematical guarantees for controlling online error criteria based either on the family-wise error rate (FWER) or the marginal false discovery rate (mFDR). We illustrate the benefit of super-uniform rewarding in real-data analyses and simulation studies. While discrete tests serve as our leading example, we also show the benefit of our method for online p-values weighting. Finally, we present extensions of our theory to online FDR control and stopped mFDR control.

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1. Introduction

1.1. Background

Multiple testing is a well-established statistical paradigm for the analysis of complex and large-scale data sets, in which each hypothesis typically corresponds to a scientific question. In the classical situation, the set of hypotheses should be pre-specified before running the statistical inference. However, in contrast to the former 'offline' setting, in many contemporary applications questions arise sequentially. A first instance of such sequential application is when testing a

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single null hypothesis repeatedly as new data are collected. For instance, this is the case for continuous monitoring of A/B tests in the information technology industry or marketing research, see Kohavi et al. (2013); Johari et al. (2019) and references therein, or Howard et al. (2021) for recent developments. A second situation is when the null hypotheses are (potentially) different and arise in a continuous stream, and accordingly decisions have to be made one at a time and prior to the termination of the stream. This is generally referred to as the online multiple testing (OMT) framework and is the focus of this paper, see, e.g., Lark (2017); Robertson et al. (2019); Kohavi et al. (2020) for application examples. This second situation can also occurs in combination with the first one to form a 'doubly-sequential' experiment (Ramdas, 2019).

1.2. Existing literature on online multiple testing

The literature aiming at controlling of various error rates in OMT has grown rapidly in the last few years. As a starting point, the family-wise error rate (FWER) is the probability of making at least one error in the past discoveries, and a typical aim is to control it at each time of the stream (for a formal definition of this and other error rates, see Section 2.2). Since controlling FWER at a given level α is a strong constraint, it requires employing a procedure that is conservative, thus generally leading to few discoveries. The typical strategy is to distribute over time the initial wealth α , e.g., performing the *i*-th test at level $\alpha \gamma_i$ for a sequence $\{\gamma_i\}_{i\geq 1}$ summing to 1. This approach is generally referred to as α -spending in the literature (Foster and Stine, 2008).

A less stringent criterion is the false discovery rate (FDR), defined as the expected proportion of false discoveries. This versatile criterion allows many more discoveries than the FWER and has known a huge success in offline multiple testing literature since its introduction by Benjamini and Hochberg (1995), both from a theoretical and practical point of view. In their seminal work on OMT, Foster and Stine (2008) extended the FDR in an online setting by considering the expected proportion of errors among the past discoveries (actually, considering rather the marginal FDR, denoted below by mFDR, which is defined as the ratio of the expectations, rather than the expectation of the ratio). The novel strategy in Foster and Stine (2008), which is called α -investing, is based on the idea that an mFDR controlling procedure is allowed to recover some α -wealth after each rejection, which slows down the natural decrease of the individual test levels. In subsequent papers, many further improvements of this method have been proposed: first, the α -investing rule has been generalized by Aharoni and Rosset (2014), while maintaining marginal FDR control. Later, Javanmard and Montanari (2018) established the (non-marginal) FDR control of these rules, including the LORD (Levels based On Recent Discovery) procedure. Then, a uniform improvement of LORD, called LORD++, has been proposed by Ramdas et al. (2017), that maintains FDR/mFDR control while extending the theory in several directions (weighting, penalties, decaying memory).

Extensions to other specific frameworks have been proposed, including rules that allow asynchronous online testing (Zrnic et al., 2021), maintain privacy

(Zhang et al., 2020), and accommodate a high-dimensional regression model (Johnson et al., 2020). Other online error criteria have also been explored, such as the false discovery exceedance (Javanmard and Montanari, 2018; Xu and Ramdas, 2021), post hoc false discovery proportion bounds (Katsevich and Ramdas, 2020), or confidence intervals with false coverage rate control (Weinstein and Ramdas, 2020).

Compared to the offline framework, the online framework has a temporal structure that needs to be taken into account, which generally leads to less powerful procedures. Hence, another important branch of the literature aims at proposing improved rules that gain more discoveries: first, following the classical 'adaptive' offline strategy, procedures can be made less conservative by implicitly estimating the amount of true null hypotheses, see the SAFFRON procedure for FDR and the adaptive-spending procedure for FWER (Ramdas et al., 2018; Tian and Ramdas, 2021). Second, under an assumption on the null distribution, increasing the number of discoveries is possible by 'discarding' tests with too large p-values (Tian and Ramdas, 2019).

A power enhancement can also be obtained by combining online procedures with other methods. A natural idea is to use more sophisticated individual tests in the first place, e.g., based on multi-armed bandits (Yang et al., 2017), or so-called 'always valid p-values', see Johari et al. (2019) and references therein. Another idea is to combine offline procedures to form 'mini-batch' rules, see Zrnic et al. (2020). Further improvements are also possible by incorporating contextual information as done by Chen and Kasiviswanathan (2020a) or using local FDR-like approach, see Gang et al. (2020). Lastly, performance boundaries have been derived by Chen and Arias-Castro (2021).

1.3. Super-uniformity

This paper considers OMT in the setting of super-uniformly distributed p-values (defined in detail in Section 2.1). Super-uniformity may originate from various sources. The first main example we have in mind, and which has been extensively investigated in the statistical literature, is super-uniformity arising from discrete p-values (described in detail in Section 5). Additionally, we show that super-uniformity can also be used in a more indirect way as a device for dealing with online p-value weighting. In the offline setting, this is a powerful and extensively studied approach, which has, however, in the online case, received little attention so far (described in detail in Section 6).

Discrete tests often originate when the tests are based on counts or contingency tables, for example:

- in clinical studies, the efficiency or safety of drugs are compared by counting patients who survive a certain period after being treated, or who experience a certain type of adverse drug reaction;
- in biology, the genotype effect on the phenotype can be tested by knocking out genes sequentially in time.

The latter example also describes the data from the International Mouse Phenotyping Consortium (IMPC, see Muñoz-Fuentes et al. (2018)), which contains many categorical variables, observed as counts and summarized in contingency tables. While this data set is frequently used (see e.g., Tian and Ramdas, 2021; Xu and Ramdas, 2021; Karp et al., 2017), the classical OMT procedures do not exploit the discrete nature of the tests, and it turns out that much more powerful procedures can be developed, see Section 5.3.

In the literature, different solutions have been proposed for dealing with the conservatism of discrete tests already at the single test level, the most straightforward one being randomization (see e.g., Habiger, 2015). While this approach possesses attractive theoretical properties, randomization is usually unacceptable in practice (Lehmann and Romano, 2022). An active research area explores discrete multiple testing in the offline setting, with the seminal works of Tarone (1990); Westfall and Wolfinger (1997); Gilbert (2005a) and the subsequent studies of Heyse (2011); Dickhaus et al. (2012); Habiger (2015); Chen et al. (2015); Döhler (2016); Chen et al. (2018); Döhler et al. (2018); Durand et al. (2019), see also references therein. The improvement achieved by such methods can be substantial and the present work shows that similar improvements are also possible in the online setting.

Finally, p-value weighting is a well-established and popular approach for improving the performance of offline multiple testing procedures. It can be traced back to Holm (1979) and has been further developed, in, e.g., Genovese et al. (2006); Wasserman and Roeder (2006); Rubin et al. (2006); Blanchard and Roquain (2008); Roquain and van de Wiel (2009); Hu et al. (2010); Zhao and Zhang (2014); Ignatiadis et al. (2016); Durand (2019); Ramdas et al. (2019) with weights that can be driven for instance by sample size, groups, or more generally by some covariates. By approaching the problem from the perspective of super-uniformity, our general method also allows seamless and flexible integration of such weighting schemes in an online context.

1.4. Contributions of the paper

In this paper, we propose uniform improvements of the classical base procedures¹ listed in Table 1, and prove control of the corresponding error rates. A distinguishing feature of our work is that we assume that a (non-trivial) upper bound for the null cumulative distribution function's (c.d.f.), called the *null bounding family*, is known (see Section 2.1). By combining this information with base procedures, we construct more efficient OMT procedures (see Table 2). The key quantity involved in this construction can be interpreted as a reward induced by the super-uniformity of the null bounding family. Figure 1 illustrates the concept in the case of discrete tests: at time t, the reward is obtained as the difference between the nominal significance level α_t and the truly achieved

¹For consistent terminology within this paper we have renamed the classical ADDIS and (the slightly modified) SAFFRON procedures (more details on the relationship of AOB to ADDIS and ALORD to SAFFRON can be found in Sections 3.3 and 4.3).

Table 1

Overview of the critical values of the base procedures for some choice of level $\alpha \in (0,1)$, adaptivity parameter $\lambda \in [0,1)$, initial wealth $W_0 \in (0,\alpha)$, and spending sequence $(\gamma_j)_{j\geq 1}$. The quantities $\mathcal{T}(\cdot)$, τ_j , $\mathcal{T}_j(\cdot)$ are given by (16), (24), (29), respectively.

Error rate	Procedure	Critical values	Results
FWER	OB AOB (ADDIS)	$\alpha_T^{\text{OB}} = \alpha \gamma_T$ $\alpha_T^{\text{AOB}} = \alpha (1 - \lambda) \gamma_{\mathcal{T}(T)}$	Tian and Ramdas (2021) Tian and Ramdas (2021)
mFDR	LORD	$\alpha_T^{\text{LORD}} = W_0 \gamma_T + (\alpha - W_0) \gamma_{T-\tau_1} + \alpha \sum_{j \ge 2} \gamma_{T-\tau_j}$	Javanmard and Montanari (2018) and Ramdas et al. (2017)
	ALORD (SAFFRON)	$\begin{split} &\alpha_T^{\text{ALORD}} = (1 - \lambda) \cdot \left(W_0 \gamma_{\mathcal{T}_0(T)} \right. \\ &\left. + (\alpha - W_0) \gamma_{\mathcal{T}_1(T)} + \alpha \sum_{j \geq 2} \gamma_{\mathcal{T}_j(T)} \right) \end{split}$	Ramdas et al. (2018) (slightly improved)

Table 2

Overview of the critical values of the rewarded procedures denoted as the corresponding base procedures, with an additional symbol " ρ " in the name. Here, α_T^{OB} , α_T^{AOB} , α_T^{LORD} , α_T^{ALORD} are the base procedures from Table 1 (with the adaptivity parameter λ defined there), ρ_t is the super-uniformity reward at time t given by (8), γ' is the SURE spending sequence defined in Section 2.4 and $\varepsilon_T = \mathbf{1}\{p_T < \lambda\}(\alpha_T - \alpha_T^0)$ is an additional adaptivity reward, for either $(\alpha_T^0, \alpha_T) = (\alpha_T^{ADB}, \alpha_T^{\rho AOB})$, or $(\alpha_T^0, \alpha_T) = (\alpha_T^{ALORD}, \alpha_T^{\rho ALORD})$, depending on the case.

Error rate	Procedure	Critical values	Results
FWER	$ ho { m OB}$	$\alpha_T^{\rho \text{OB}} = \alpha_T^{\text{OB}} + \sum_{t=1}^{T-1} \gamma_{T-t}' \rho_t$	Theorem 3.1
	$ ho { m AOB}$	$\alpha_T^{\rho \text{AOB}} = \alpha_T^{\text{AOB}} + \sum_{\substack{1 \le t \le T-1 \\ p_t > \lambda}} \gamma'_{T-t} \rho_t + \varepsilon_{T-1}$	Theorem 3.2
		$\alpha_T^{\rho \text{LORD}} = \alpha_T^{\text{LORD}} + \sum_{t=1}^{T-1} \gamma_{T-t}' \rho_t$ $\alpha_T^{\rho \text{ALORD}} = \alpha_T^{\text{ALORD}} + \sum_{t=1}^{T-1} \gamma_{T-t}' \rho_t + \varepsilon_{T-1}$	Theorem 4.1 Theorem 4.2
		$\substack{1 \leq t \leq T-1 \\ p_t > \lambda}$	

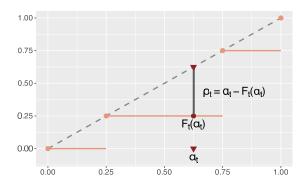


FIG 1. Super-uniformity reward ρ_t at time t (length of the vertical line) as defined by (8) (see Section 2.3) for a given discrete bounding function F_t (orange step function) and a critical value α_t (triangle). The dashed line is the identity function $x \in [0,1] \mapsto x$.

significance level $F_t(\alpha_t)$. We use the acronym SUR (Super-Uniform-Reward) to refer to these new procedures. When we use the uniform null bounding family (i.e., in the classical framework), our SUR procedures reduce to their base counterparts. Our main contributions are as follows:

- We propose two new SUR procedures for online FWER control in Section 3: the first one (ρ OB) uniformly improves upon the Online Bonferroni procedure (OB), while the second (ρ AOB) uniformly improves upon the adaptive spending procedure of Tian and Ramdas (2021) (AOB).
- We propose two new SUR procedures for online mFDR control in Section 4: the first one (ρLORD) uniformly improves upon the LORD++ procedures of Javanmard and Montanari (2018); Ramdas et al. (2017) (LORD), while the second one (ρALORD) uniformly improves upon the SAFFRON procedure of Ramdas et al. (2018) (ALORD).
- We present a general and simple way of constructing SUR procedures for any base procedure satisfying some mild conditions, see Section 3.4 for FWER and Section 4.4 for mFDR. This allows us to obtain concise proofs for all our results, which are deferred to the supplement, see Appendix 8.
- Application to discrete data: we evaluate the performances of the new SUR procedures on discrete data with simulated experiments (Section 5.2) and for a classical real data set (Section 5.3), where each hypothesis is tested using a (discrete) Fisher exact test. The gain in power is shown to be substantial.
- Application to online p-value weighting: our new SUR procedures can be used to derive weighted online FWER and mFDR controlling procedures. The p-value weighting is carried out by rescaling the 'raw' weights in a certain way so that the weighted p-value distributions become super-uniform and our methodology can be applied. The new online procedures are shown to outperform existing ones both on simulated and real data (Section 6).
- We show that our mFDR-controlling online methods also control the mFDR at any stopping time in Appendix 9, and we provide a modification of ρ LORD that controls the (non marginal) FDR in Appendix 10.

For easier readability of the paper, a succinct overview of our work is presented in Tables 1 and 2. It lists the base and SUR procedures and provides links to definitions and results for error rate control. All our numerical experiments (simulations and application) are reproducible from the code provided in the repository https://github.com/iqm15/SUREOMT.

Finally, the appendices contain further material including proofs, alternative approaches and additional numerical studies.

1.5. Relation to adaptive discarding

As Tian and Ramdas (2019) point out, online multiple testing procedures frequently suffer from significant power loss if the null *p*-values are too conservative. In Tian and Ramdas (2021) (FWER control) and Tian and Ramdas (2019)

(mFDR control), the authors propose adaptive discarding (ADDIS) approaches as improved methods. In particular, an idea is to use a discarding rule, that avoids testing a null when the corresponding p-value exceeds a given threshold. For the particular type of super-uniformity induced by discrete tests, we show that the discarding rule is less efficient than the SUR method, at least in the settings of Sections 5.2 and 5.3.

2. Preliminaries

2.1. Setting, procedure and assumptions

Let $X = (X_t, t \in \{1, 2, \dots\})$ be a process composed of random variables. We denote the distribution of X by P, which is assumed to belong to some distribution set \mathcal{P} . We consider an online testing problem where, at each time $t \geq 1$, the user only observes variable X_t and should test a new null hypothesis H_t , which corresponds to some subset of \mathcal{P} . We let $\mathcal{H}_0 = \mathcal{H}_0(P) = \{t \geq$ 1: H_t is satisfied by P} the set of (unknown) times where the corresponding null hypothesis is true. Throughout the manuscript, we focus on decisions based upon p-values. Hence, we suppose that at each time t, we have at hand a pvalue $p_t = p_t(X) \in [0,1]^2$ (typically depending only on X_t although this is not necessary) for testing H_t , and we consider online multiple testing procedures based on p-value thresholding. This means that each null H_t is rejected whenever $p_t(X) \leq \alpha_t$, where $\alpha_t \in [0, \infty)$ is a nonnegative threshold, called a critical value, that is allowed to depend on the past decisions. More precisely, we denote $R_t = \mathbf{1}\{p_t(X) \leq \alpha_t\}, C_t = \mathbf{1}\{p_t(X) \geq \lambda\}$ for all $t \geq 1$ and assume that each α_t is measurable with respect to the σ -field $\mathcal{F}_{t-1} = \sigma(R_1, \ldots, R_{t-1}, C_1, \ldots, C_{t-1})$. Here, $\lambda \in [0,1]$ is a parameter that is used for designing adaptive procedures. The non-adaptive case is recovered as a special case by setting $\lambda = 0$, in which case $\mathcal{F}_{t-1} = \sigma(R_1, ..., R_{t-1})$.

In the literature, this property is referred to as predictability, see Ramdas et al. (2017). Throughout the manuscript, an online multiple testing procedure is identified with a family $\mathcal{A} = \{\alpha_t, t \geq 1\}$ of such predictable critical values. Let us now state the assumptions used in what follows. First, recall the classical super-uniformity assumption:

$$\mathbf{P}_{X \sim P}(p_t(X) \le u) \le u \text{ for all } u \in [0, 1], \text{ and } P \in \mathcal{P} \text{ with } t \in \mathcal{H}_0,$$
 (1)

which means that each test rejecting $H_{0,t}$ when $p_t(X)$ is smaller than or equal to u is of level u. Here, we typically consider a setting where these tests may have a more stringent level. Formally, we assume that at each time t, a known null bounding function $F_t: [0,1] \to [0,1]$ is available, satisfying

$$\mathbf{P}_{X \sim P} (p_t(X) \le u) \le F_t(u) \le u, \text{ for all } u \in [0, 1], \text{ and } P \in \mathcal{P} \text{ with } t \in \mathcal{H}_0.$$
(2)

 $^{^{2}}$ In the sequel, we write p_{t} instead of $p_{t}(X)$ when there is no ambiguity.

Note that we will sometimes also consider $F_t(u)$ for $u \ge 1$, in which it is to be understood as $F_t(u \land 1)$. The family $\mathfrak{F} = \{F_t, t \ge 1\}$ of null-bounding functions will be referred to as the *null bounding family*. Note that (2) reduces to (1) when choosing $F_t(u) = u$ for all u, but encompasses other cases by choosing differently the null bounding family. Typically, for discrete tests, it is well-known that $F_t(u)$ can be (much) smaller than u, see Example 2.1 for more details. Second, another important assumption is the *online independence* within the p-value process:

$$p_t(X)$$
 is independent of the past decisions \mathcal{F}_{t-1} for all $t \in \mathcal{H}_0$ and $P \in \mathcal{P}$.

Assumption (3) holds for instance, in the case where $p_t(X)$ only depends on X_t and the variables in X are all mutually independent, which means that the data are collected independently at each time.

Remark 2.1. In this manuscript, results are often based on assumptions (2) and (3). In all these results, these two assumptions can be replaced by the weaker condition

$$\forall u \in [0, 1], \forall t \in \mathcal{H}_0 \text{ and } P \in \mathcal{P}, \mathbf{P}_{X \sim P}(p_t(X) \le u | \mathcal{F}_{t-1}) \le F_t(u) \le u \text{ a.s.}$$
 (4)

When choosing the null bounding family $F_t(u) = u$ for all u, the latter condition is sometimes referred to as SuperCoAD (super-uniformity conditionally on all discoveries), see Ramdas et al. (2017).

Throughout the paper, we investigate the two following prototypical examples of super-uniformity.

Example 2.1. Our leading example is the case where a discrete test statistic is used for inference in each individual test. Typical instances include tests for analyzing counts such as Poisson and Fisher's exact test, but also permutation tests (Romano and Wolf, 2005), see Appendix 13 for more details on these discrete tests. In discrete testing, each p-value $p_t(X)$ has its own support S_t (known and not depending on P), that is a finite set (or, in full generality, a countable set with 0 or 1 as the only possible accumulation points). A null bounding family satisfying (2) can easily be derived by considering F_t , the right-continuous step function that jumps at each point of S_t , see Figure 1 as an example. The support S_t can depend on t so that discrete testing can also induce heterogeneity over time. In Section 5 we apply our SUR approach to discrete tests.

Example 2.2. Our secondary example is p-value weighting, where we start from continuous p-values that are uniform under the null (i.e. corresponding to the "usual" case). These p-values are then weighted using external information in order to increase power. Our SUR approach is developed for weighted p-values in Section 6.

2.2. Error rates and power

Let us define the criteria that we use to measure the quality of a given procedure $\mathcal{A} = \{\alpha_t, t \geq 1\}$. For each $T \geq 1$, let $\mathcal{R}(T) = \{t \in \{1, ..., T\} : p_t(X) \leq \alpha_t\}$

denote the set of rejection times of the procedure A, up to time T. We consider the two following classical online criteria for type I error rates:

$$FWER(\mathcal{A}, P) := \sup_{T \geq 1} \{FWER(T, \mathcal{A}, P)\};$$

$$FWER(T, \mathcal{A}, P) := \mathbf{P}_{X \sim P} (|\mathcal{H}_0 \cap \mathcal{R}(T)| \geq 1);$$

$$mFDR(\mathcal{A}, P) := \sup_{T \geq 1} \{mFDR(T, \mathcal{A}, P)\};$$

$$mFDR(T, \mathcal{A}, P) := \frac{\mathbf{E}_{X \sim P}(|\mathcal{H}_0 \cap \mathcal{R}(T)|)}{\mathbf{E}_{X \sim P}(1 \vee |\mathcal{R}(T)|)},$$

$$(6)$$

with the convention 0/0 = 0. In words, when controlling the online FWER at level α , one has the guarantee that, at each fixed time T, the probability of making at least one false discovery before time T is below α . Since FWER control does not tolerate any false discovery (with high probability), it is generally considered a stringent criterion. By contrast, when controlling the online mFDR, at each time T, the expected number of false discoveries before time T need not be small provided the expected number of discoveries is large enough. While online FWER has been investigated in Tian and Ramdas (2021), online mFDR control is generally less conservative (that is, allows more discoveries), and is widely used in an online context, see Foster and Stine (2008); Ramdas et al. (2017, 2018). The false discovery rate (FDR) is closely related to the mFDR: it is defined by using the expectation of the ratio, instead of the ratio of the expectations as in (6). Controlling the FDR generally requires more assumptions, while mFDR is particularly useful in an online context (we refer the reader to Section 1.1 of Zrnic et al. (2021) for more discussions on this). For a given error rate, we aim at deriving procedures that maximize power. For any procedure A, we define the power as the expected proportion of signal the procedure can detect, that is,

$$Power(T, \mathcal{A}, P) := \frac{\mathbf{E}_{X \sim P} (|\mathcal{H}_1 \cap \mathcal{R}(T)|)}{1 \vee |\mathcal{H}_1|}, \tag{7}$$

where \mathcal{H}_1 is the set of times of false nulls, that is, the complement of \mathcal{H}_0 in $\{1, 2, \dots\}.$

While this power notion will be used in our numerical experiments to compare procedures, our theoretical results will use a stricter comparison criterion. For two online procedures $\mathcal{A} = \{\alpha_t, t \geq 1\}$ and $\mathcal{A}' = \{\alpha_t', t \geq 1\}$, we say that \mathcal{A}' uniformly dominates A when $\alpha'_t \geq \alpha_t$ for all $t \geq 1$ (almost surely). This implies that, almost surely, \mathcal{A}' makes more discoveries than \mathcal{A} , in the sense that the set of discoveries of \mathcal{A} is contained in the one of \mathcal{A}' , that is, $\mathcal{R}(T) \subset \mathcal{R}'(T)$ for all T > 1 (a.s.). In particular, this implies the same domination for the true discovery sets and thus in particular $\operatorname{Power}(T, \mathcal{A}, P) \leq \operatorname{Power}(T, \mathcal{A}', P)$ for all T > 1. With this terminology, we can restate the aim of this work as follows: construct valid OMT procedures that uniformly dominate their base procedures by incorporating the null bounding family function F_t given in (2).

Remark 2.2. There is no consensus regarding the most adequate definition of power in online testing literature. The concept of uniform domination that we

use in this paper is much stronger than, e.g., the asymptotic power considered by Javanmard and Montanari (2018). It may, however, not be particularly appropriate if the base procedure $\mathcal A$ is chosen poorly. Since the base procedures given in Table 1 are standard in our setting, the domination criterion seems to be reasonable.

2.3. Wealth and super-uniformity reward

In the Generalized Alpha-Investing (GAI) paradigm (see Xu and Ramdas (2021) and the references given therein), the nominal level α , at which one wants to control the type I error rate, can be seen as an overall error budget – or wealth – that may be spent on testing hypotheses in the course of an online experiment. For a given OMT procedure \mathcal{A} , it is possible to define a suitable wealth function $W(T) = W(T, \mathcal{A}, P)$, such that W(T) represents the wealth available at time T for further testing. As a case in point, Xu and Ramdas (2021) define the (nominal) wealth function for the online Bonferroni procedure by $W^{\text{nom}}(T) = \alpha - \sum_{t=1}^{T} \alpha \gamma_t$. Generalizing this expression for arbitrary null distributions we obtain the 'true' or 'effective' wealth $W^{\text{eff}}(T) = \alpha - \sum_{t=1}^{T} F_t(\alpha \gamma_t)$, where F_t is a null-bounding function. In the super-uniform setting, assumption (2) implies $W^{\text{nom}}(T) \leq W^{\text{eff}}(T)$, and as the two orange curves in Figure 2 illustrate, the discrepancy can be quite large.

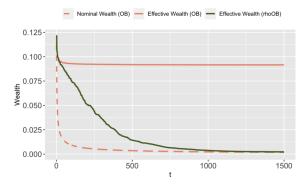


FIG 2. Nominal wealth for OB (dashed orange curve, 'what we want to spend'), effective wealth for OB (solid orange curve, 'what OB actually spends') and effective wealth for ρ OB (solid green curve, 'what ρ OB actually spends') for the male mice from the IMPC data (see Section 5.3 for more details).

While the user thinks the procedure is spending the budget over time according to the nominal wealth given by the dashed orange curve, in reality, the procedure is under-utilizing wealth, as the solid orange true wealth curve indicates. This unnecessarily austere spending behaviour makes the online Bonferroni procedure sub-optimal. In addition, this phenomenon extends to the other procedures and error rates listed in Table 1 as well. Our proposed solution incorporates super-uniformity so that its wealth function behaves more like the targeted nominal wealth, as depicted by the green curve in Figure 2.

For incorporating super-uniformity, we introduce the super-uniformity reward (SUR), a key quantity in our work. For any procedure $\mathcal{A} = \{\alpha_t, t \geq 1\}$ and null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$, the super-uniformity reward ρ_t at time t is defined by

$$\rho_t = \rho_t(\alpha_t, F_t) := \alpha_t - F_t(\alpha_t), \quad t \ge 1.$$
(8)

Note that (2) always implies $\rho_t \geq 0$ for all $t \geq 1$. In the case of discrete testing (Example 2.1), we have $F_t(\alpha_t) = 0$ when α_t is below the infimum of the support \mathcal{S}_t . This produces the maximum possible super-uniformity reward at time t, that is, $\rho_t = \alpha_t$. Conversely, when $\alpha_t \in \mathcal{S}_t$, we have $F_t(\alpha_t) = \alpha_t$ and we have no super-uniformity reward at time t, that is, $\rho_t = 0$. In general, we have $\rho_t \in [0, \alpha_t]$, its actual value depending on the discreteness of the test (that is on the steps of F_t) and of the value of α_t . The super-uniformity reward is illustrated in Figure 1 for a single distribution F_t and value α_t . Mathematically, ρ_t is simply the difference between the nominal significance level α_t and the truly achieved significance level $F_t(\alpha_t)$. In terms of wealth, ρ_t can be interpreted as the fraction of nominal significance level which the OMT procedure was unable to 'spend' due to super-uniformity. Intuitively, it seems clear that this amount can be put aside and be re-allocated to the subsequent tests to increase the future critical values $(\alpha_T, T \ge t + 1)$. In Sections 3 and 4, we show in detail how this can be done without sacrificing type I error control.

2.4. Spending sequences

As Table 1 displays, the base procedures we use are parametrized by a sequence $\gamma = (\gamma_t)_{t\geq 1}$ of non-negative values, such that $\sum_{t\geq 1} \gamma_t \leq 1$, which we refer to as the spending sequence. The spending sequence controls the rate at which the wealth is spent in the course of the online experiment (for instance, see (10) for the online Bonferroni procedure). However, finding suitable spending sequences is not trivial: there is a trade-off between saving wealth for large values of T and the ability to make discoveries in the not-too-distant future. Typical choices for γ in the literature are:

- $\gamma_t \propto t^{-q}$ for all t for some q > 1, see Tian and Ramdas (2021);
- $\gamma_t \propto (t+1)^{-1} \log^{-q}(t+1)$ for all t, for some q > 1, see Tian and Ramdas

Throughout the paper, we choose $\gamma_t \propto t^{-q}$ with q=1.6, as suggested by previous literature. In the base procedures listed in Table 1, there are two potential sources of wealth: the initial wealth invested at T=0, and the rejection reward that can be earned by rejections for investing procedures (i.e., mFDR controlling procedures). When one can use super-uniformity reward as described in Section 2.3, an additional source of wealth comes into play. Indeed, our approach uses an additional SUR spending sequence γ' to smoothly incorporate all the rewards collected up to time T to compute the new critical value α_T . This SUR spending sequence could be chosen for instance from one of the smoothing sequences listed above. Here, we focus on the following choice:

$$\gamma_t' = \gamma_t'(h) = \mathbf{1}\{t \le h\}/h, \quad t \ge 1,$$
 (9)

where $h \geq 1$ is a suitably chosen integer. Since this leads to procedures that spread rewards uniformly over a finite horizon of length h, we refer to (9) – by analogy with non-parametric density estimation – as a rectangular kernel with bandwidth h. Finally, another idea introduced by Ramdas et al. (2018); Tian and Ramdas (2021) in order to slow down the natural decay in the α_t sequence is to consider $\gamma_{\mathcal{T}(t)}$ where $\mathcal{T}(t)$ is a slowed down clock, see (16) and (29) below. As we will see in Section 3.3 and Section 4.3, this technique can also be combined with a suitable super-uniformity reward.

3. Online FWER control

In this section, we aim at finding procedures \mathcal{A} such that FWER $(\mathcal{A}, P) \leq \alpha$ for some targeted level $\alpha \in (0,1)$. We begin with a simple presentation of our approach to improve the online Bonferroni procedure with a 'greedy' super-uniformity reward, and then turn to a smoother spending of the super-uniformity reward (Theorem 3.1). This approach is then applied in combination with the adaptive online procedure introduced by Tian and Ramdas (2021) (Theorem 3.2). Finally, a general result is provided (Theorem 3.3) that allows to reward any procedure controlling the online FWER in some specific way. This allows unifying all results obtained in this section while further extending the scope of our methodology.

3.1. Warming-up: online Bonferroni procedure and a first greedy reward

For any given spending sequence $\gamma = (\gamma_t)_{t \geq 1}$, a well-known online FWER controlling procedure is the online Bonferroni procedure, $\mathcal{A}^{\text{OB}} = \mathcal{A}^{\text{OB}}(\alpha, \gamma) := \{\alpha_t^{\text{OB}}, t \geq 1\}$, defined by

$$\alpha_T^{\text{OB}} := \alpha \gamma_T, \quad T \ge 1.$$
 (10)

It is also called Alpha-Spending rule (Foster and Stine, 2008) in the context of online FWER control, see Tian and Ramdas (2021). It is straightforward to check that \mathcal{A}^{OB} controls the FWER under the classical super-uniformity condition (1): by the Markov inequality, for all $T \geq 1$,

$$\text{FWER}(T, \mathcal{A}^{\text{OB}}, P) \le \mathbf{E}_{X \sim P} \Big(\sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_0, p_t \le \alpha \gamma_t \} \Big)$$
 (11)

$$\leq \sum_{t \in \mathcal{H}_0} \mathbf{P}_{X \sim P}(p_t \leq \alpha \gamma_t) \leq \sum_{t \in \mathcal{H}_0} \alpha \gamma_t \leq \alpha.$$
 (12)

Let us now present the rationale behind our approach in this simple case. Assume more generally that we have at hand a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2). For any procedure $\mathcal{A} = \{\alpha_t, t \geq 1\}$ (with deterministic α_t) the above reasoning leads to the following upper bound on the FWER:

$$FWER(T, \mathcal{A}, P) \le \alpha_T + \sum_{t=1}^{T-1} F_t(\alpha_t) =: \widehat{FWER}(T, \mathcal{A}).$$
 (13)

Thus, FWER is controlled if we can guarantee that $\widetilde{\text{FWER}}(T, \mathcal{A}) \leq \alpha$ for all $T \in \{1, 2, \ldots\}$. There are various ways of achieving this. A simple solution that takes the spending schedule of \mathcal{A}^{OB} as a benchmark, consists of choosing $\alpha_T = \sum_{t=1}^T \alpha \gamma_t - \sum_{t=1}^{T-1} F_t(\alpha_t)$. The latter is a recursive relation that allows us to define a new procedure $\mathcal{A} = \{\alpha_t, t \geq 1\}$ controlling the FWER. Indeed, since $\alpha_1 = \alpha \gamma_1$ and for $T \geq 2$, $\alpha_T - \alpha_{T-1} = \alpha \gamma_T - F_{T-1}(\alpha_{T-1})$, this leads to the simple rule

$$\alpha_T = \alpha \gamma_T + \rho_{T-1}, \quad T \ge 1, \tag{14}$$

where $\rho_{T-1} = \alpha_{T-1} - F_{T-1}(\alpha_{T-1})$ is the super-uniformity reward (8) at time T-1 (with the convention $\rho_0 = 0$). In addition, from (2), we have $\rho_{T-1} \geq 0$, and the critical values (14) uniformly dominate the online Bonferroni critical values (10) (the obtained critical values are in particular nonnegative, thus defining a valid OMT procedure). The algorithm behind critical values (14) can be described as 'greedy', because it spends the complete super-uniformity reward ρ_{T-1} obtained at step T-1 for increasing the next critical value α_T .

3.2. Smoothing out the super-uniformity reward

The greedy policy described in the previous section is not always appropriate when time is considered on a potentially large scale, because the sequence of critical values might decrease too quickly. Instead, we can smooth this effect over time, by distributing the reward collected at time T-1 over all times following T. To formalize this idea, we introduce a SUR spending sequence (see also Section 2.4), which is defined as a non-negative sequence $\gamma' = (\gamma'_t)_{t\geq 1}$ such that $\sum_{t\geq 1} \gamma'_t \leq 1$. While this definition is mathematically the same as the definition of a spending sequence, the role of the SUR spending sequence is different, so we use a different name for it.

Definition 3.1. For any spending sequence γ and any SUR spending sequence γ' , the online Bonferroni procedure with super-uniformity reward, denoted by $\mathcal{A}^{\rho OB} = \{\alpha_t^{\rho OB}, t \geq 1\}$, is defined by the recursion

$$\alpha_T^{\rho OB} = \alpha \gamma_T + \sum_{t=1}^{T-1} \gamma'_{T-t} \rho_t, \quad T \ge 1, \tag{15}$$

where $\rho_t = \alpha_t^{\rho OB} - F_t(\alpha_t^{\rho OB})$ denotes the super-uniformity reward at time t for that procedure.

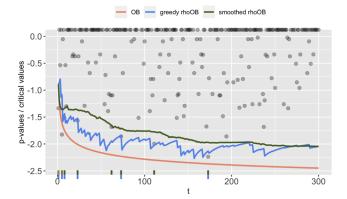


FIG 3. Sequences of critical values for Bonferroni procedures with different rewards over time $1 \le t \le T = 300$ (simulated data): base Bonferroni critical values (10) (orange line), rewarded with the greedy approach (14) (blue line), and with the rectangular kernel SUR spending sequence (15) (h = 10, green line). The rug plots display the time of discoveries for each procedure with the corresponding color. The Y-axis has been transformed by $y \mapsto -\log(-\log(y))$. The grey dots denote the p-value sequence (those equal to 1 are displayed at the top of the picture). The spending sequence is $\gamma_t \propto t^{-1.6}$.

Note that taking $\gamma'=(1,0,\ldots 0)$ recovers the 'greedy' critical values (14). For the rectangular kernel SUR spending sequence given by (9), we have $\sum_{t=1}^{T-1} \gamma'_{T-t} \rho_t = h^{-1} \sum_{t=1 \vee (T-h)}^{T-1} \rho_t$, which we interpret as a uniform spending of the super-uniformity reward over the last h time points. As shown in Figure 3, the corresponding sequence of critical values (green line) is more 'stable' than the one using the greedy approach (blue line), allowing for some additional discoveries (on this simulated data).

The following result provides FWER control of the new rewarded critical values (15), for a general SUR spending sequence.

Theorem 3.1. Consider the setting of Section 2.1, where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. For any spending sequence γ and any SUR spending sequence γ' , consider the online Bonferroni procedure $\mathcal{A}^{OB} = \{\alpha_t^{OB}, t \geq 1\}$ (10), and the online Bonferroni with super-uniformity rewards $\mathcal{A}^{POB} = \{\alpha_t^{POB}, t \geq 1\}$ (15). Then we have $\mathrm{FWER}(\mathcal{A}^{POB}, P) \leq \alpha$ for all $P \in \mathcal{P}$, while \mathcal{A}^{POB} uniformly dominates \mathcal{A}^{OB} .

This result will be a consequence of a more general result, see Section 3.4.

3.3. Rewarded adaptive online Bonferroni

It is apparent from (11)-(12) that there is some looseness when upper-bounding $\sum_{t\in\mathcal{H}_0}\gamma_t$ by $\sum_{t\geq 1}\gamma_t$ which may lead to unnecessarily conservative procedures. We may attempt to avoid this loss in efficiency by considering a spending sequence γ satisfying the condition $\sum_{t\in\mathcal{H}_0}\gamma_t\leq 1$ which is more liberal than $\sum_{t\geq 1}\gamma_t\leq 1$. In words, this means that the index t in the sequence $\{\gamma_t,t\geq 1\}$

should only be incremented when we are testing an hypothesis H_t with $t \in \mathcal{H}_0$. Since \mathcal{H}_0 is unknown, such a modification cannot be implemented directly in the γ sequence. Nevertheless, the ADDIS procedure proposed by Tian and Ramdas (2021) works by replacing the unknown set \mathcal{H}_0 by an estimate $\{1\} \cup \{t \geq 2 : p_{t-1} \geq \lambda\}$ for some parameter $\lambda \in (0,1)$, and to correct the introduced error in the thresholds α_t to maintain the FWER control. More formally, we follow Tian and Ramdas (2021) by introducing the re-indexation functional $\mathcal{T}: \{1,\ldots\} \to \{1,\ldots\}$ defined by

$$\mathcal{T}(T) = 1 + \sum_{t=2}^{T} \mathbf{1}\{p_{t-1} \ge \lambda\}, \quad T \ge 1.$$
 (16)

Since a large p-value is more likely to be linked to a true null, $\mathcal{T}(T)$ is used to account for the number of true nulls before time T (note that this estimate is nevertheless biased). From an intuitive point of view, $\mathcal{T}(T)$ slows down the time by only incrementing time when the preceding p-value is large enough. This idea leads to the adaptive online Bonferroni procedure introduced by Tian and Ramdas (2021) (called 'Adaptive spending' there), with spending sequence γ and adaptivity parameter $\lambda \in [0,1)$, denoted here by $\mathcal{A}^{\text{AOB}} = \{\alpha_t^{\text{AOB}}, t \geq 1\}$, and given by

$$\alpha_T^{\text{AOB}} = \alpha (1 - \lambda) \gamma_{\mathcal{T}(T)}, \quad T \ge 1.$$
 (17)

It recovers the standard online Bonferroni procedure when $\lambda=0$ (because $\mathcal{T}(T)=T$ for $T\geq 1$ in that case), but leads to different thresholds when $\lambda>0$. Comparing \mathcal{A}^{AOB} to \mathcal{A}^{OB} , no procedure uniformly dominates the other. An improvement of \mathcal{A}^{AOB} over \mathcal{A}^{OB} is expected to hold when there are many false null hypotheses in the data, and increasingly so if the signal occurs early in the time sequence, see the numerical experiments in Section 5.2. In addition, note that the critical value α_T^{AOB} depends on the data X_1,\ldots,X_{T-1} and thus is random. As a result, the adaptive approach requires additional distributional assumptions compared with the online Bonferroni procedure. In Tian and Ramdas (2021), \mathcal{A}^{AOB} is proved to control the FWER under (1) and (3) (actually under the slightly more general condition (4) with F_t equal to identity). Let us now use this approach in combination with the super-uniformity reward.

Definition 3.2. For any spending sequence γ , any SUR spending sequence γ' , and $\lambda \in [0,1)$, the adaptive online Bonferroni procedure with super-uniformity reward, denoted by $\mathcal{A}^{\rho AOB} = \{\alpha_t^{\rho AOB}, t \geq 1\}$, is defined by

$$\alpha_T^{\rho_{AOB}} = \alpha (1 - \lambda) \gamma_{\mathcal{T}(T)} + \sum_{\substack{1 \le t \le T - 1 \\ p_t \ge \lambda}} \gamma'_{T-t} \rho_t + \varepsilon_{T-1}, \quad T \ge 1, \tag{18}$$

where $\rho_t = \alpha_t^{\rho^{AOB}} - F_t(\alpha_t^{\rho^{AOB}})$ denotes the super-uniformity reward at a time t, and $\varepsilon_{T-1} = \mathbf{1}\{p_{T-1} < \lambda\}(\alpha_{T-1} - \alpha(1-\lambda)\gamma_{T(T-1)})$ is an additional 'adaptive' reward (convention $\varepsilon_0 = 0$).

³The so-called 'discarding' part of the method proposed by Tian and Ramdas (2021) cannot be implemented in our setting because the F_t are not convex, as discussed in Section 1.5.

This class of procedures reduces to the class of procedures (15) introduced in the previous section by setting $\lambda=0$. However, when $\lambda>0$ the class is different since the term $\alpha(1-\lambda)\gamma_{T(T)}$, which comes from α_T^{AOB} , makes the threshold random. Also, the super-uniformity reward is only collected at time $t\leq T-1$ where $p_t\geq \lambda$. The latter is well expected from the motivation of the adaptive approach described above: when $p_t<\lambda$, no testing is performed so no reward could be obtained from ρ_t . Nevertheless, note that the additional term ε_{T-1} allows to collect some reward at time T-1 in the case where $p_{T-1}<\lambda$. Since this term only appears in critical values of adaptive procedures, we call it the 'adaptive' reward. It is linked to the super-uniformity reward in that no adaptive reward can be obtained if no super-uniformity reward has been collected in the past. The following result shows that this approach is valid from the perspective of FWER control.

Theorem 3.2. Consider the setting of Section 2.1 where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. For any spending sequence γ , any SUR spending sequence γ' and $\lambda \in [0,1)$, consider the adaptive online Bonferroni procedure $\mathcal{A}^{AOB} = \{\alpha_t^{AOB}, t \geq 1\}$ (17) and the adaptive online Bonferroni with super-uniformity rewards $\mathcal{A}^{\rho AOB} = \{\alpha_t^{\rho AOB}, t \geq 1\}$ (18). Then, assuming that the model \mathcal{P} is such that (3) holds, we have $\mathrm{FWER}(\mathcal{A}^{\rho AOB}, P) \leq \alpha$ for all $P \in \mathcal{P}$, while $\mathcal{A}^{\rho AOB}$ uniformly dominates \mathcal{A}^{AOB} .

Theorem 3.2 relies on a more general result (Theorem 3.3 below). Note that, contrary to Theorem 3.1, Theorem 3.2 needs an independence assumption, which is due to the adaptive methodology that makes the critical values random. If this independence assumption holds, we show in Section 5.2 that $\mathcal{A}^{\rho AOB}$ can indeed improve $\mathcal{A}^{\rho OB}$, while it always improves the procedure \mathcal{A}^{AOB} of Tian and Ramdas (2021) (as guaranteed by the above theorem).

3.4. Rewarded version for base FWER controlling procedures

In this section we present a general result stating that any procedure ensuring online FWER control (in a specific way) can be rewarded using super-uniformity while maintaining the FWER control. First, we can extend (13) by defining an adaptive estimator

$$\widehat{\text{FWER}}_{\lambda}(T, \mathcal{A}) = \frac{\alpha_T + \sum_{\substack{1 \le t \le T-1, \\ p_t \ge \lambda}} F_t(\alpha_t)}{1 - \lambda}.$$
 (19)

The following lemma holds as a direct consequence of Lemma 8.1.

Lemma 3.1. Assuming that (2) holds, for any procedure $A = (\alpha_t, t \ge 1)$, we have for all $\lambda \in [0, 1)$,

$$FWER(T, \mathcal{A}, P) \le \mathbf{E}_{X \sim P} \Big[\widehat{FWER}_{\lambda}(T, \mathcal{A}) \Big], \tag{20}$$

if either (3) holds or if the critical values $(\alpha_t, t \geq 1)$ are deterministic.

This gives rise to the following theorem.

Theorem 3.3. Assuming that (2) holds, consider any procedure $A^0 = (\alpha_t^0, t \ge 1)$ satisfying almost surely, for some $\lambda \in [0, 1)$ and for all $T \ge 1$,

$$\alpha_T^0 + \sum_{\substack{1 \le t \le T^{-1}, \\ n_t > \lambda}} \alpha_t^0 \le (1 - \lambda)\alpha. \tag{21}$$

Then the following holds:

- (i) \mathcal{A}^0 controls the online FWER, that is, FWER(\mathcal{A}^0, P) $\leq \alpha$ for all $P \in \mathcal{P}$, either if the α_T^0 are deterministic for all $T \geq 1$, or if (3) holds;
- (ii) for any SUR spending sequence $\gamma' = (\gamma'_t, t \ge 1)$, the procedure $\mathcal{A} = (\alpha_t, t \ge 1)$, corresponding to the rewarded \mathcal{A}^0 , and defined by, for all $T \ge 1$,

$$\alpha_{T} = \alpha_{T}^{0} + \sum_{\substack{1 \le t \le T-1 \\ p_{t} \ge \lambda}} \gamma_{T-t}'(\alpha_{t} - F_{t}(\alpha_{t})) + \mathbf{1}\{p_{T-1} < \lambda\}(\alpha_{T-1} - \alpha_{T-1}^{0}),$$
(22)

controls the online FWER, that is, $FWER(A, P) \leq \alpha$ for all $P \in \mathcal{P}$, either if the α_T are deterministic for all $T \geq 1$, or if (3) holds.

Theorem 3.3 is proved in Section 8.1. Condition (21) is essentially the same as Condition (20) derived in Tian and Ramdas (2021). It is satisfied by the online Bonferroni procedure ($\mathcal{A}^0 = \mathcal{A}^{\text{OB}}$), and the online adaptive Bonferroni procedure ($\mathcal{A}^0 = \mathcal{A}^{\text{AOB}}$). While this is obvious for \mathcal{A}^{OB} , the case of \mathcal{A}^{AOB} requires to carefully check how the functional $\mathcal{T}(\cdot)$ (16) slows down the time, which is done in Lemma 8.3. Statement (i) of Theorem 3.3 thus states the online FWER control for these procedures. Statement (ii) of Theorem 3.3 is our main contribution and reduces to Theorems 3.1 and 3.2, when choosing $\mathcal{A}^0 = \mathcal{A}^{\text{OB}}$ and $\mathcal{A}^0 = \mathcal{A}^{\text{AOB}}$, respectively. This recovers the rewarded procedures $\mathcal{A}^{\rho \text{OB}}$ and $\mathcal{A}^{\rho \text{AOB}}$ discussed in the previous sections: compare (22) to (15) (with $\lambda = 0$), and (22) to (18). Nevertheless, other choices for \mathcal{A}^0 satisfying (21) are possible. According to our general result, any such choice is compatible with our reward methodology.

4. Online mFDR control

In this section, we aim at finding procedures \mathcal{A} such that mFDR(\mathcal{A}, P) $\leq \alpha$ for some targeted level $\alpha \in (0,1)$. We follow the same route as for the FWER: we start with a presentation of the super-uniformity reward to the classical LORD++ procedure (Ramdas et al., 2017, called just LORD hereafter for short), and then turn to adaptive counterparts.

Finally, we propose a general result encompassing all these cases. In this section, we follow the notation of Ramdas et al. (2017) for online mFDR control. For any procedure $\mathcal{A} = \{\alpha_t, t \geq 1\}$ and realization of the *p*-value process, let us

denote

$$R(T) = \sum_{t=1}^{T} R_t, \quad R_t = \mathbf{1}\{p_t(X) \le \alpha_t\},$$
 (23)

the number of rejections of the procedure up to time T, and

$$\tau_j = \min\{t \ge 1 : R(t) \ge j\} \quad (\tau_j = +\infty \text{ if the set is empty}),$$
 (24)

the first time that the procedure makes j rejections, for any $j \geq 1$.

Finally, we define the following additional notation: the critical values are defined recursively by involving a suitable functional $\mathbf{A}_t(R_1,\ldots,R_{t-1})$ (Sections 4.1 and 4.2) or $\mathbf{A}_t(R_1,\ldots,R_{t-1},C_1,\ldots,C_{t-1})$ (Sections 4.3 and 4.4), where the sequence R_1,\ldots,R_{t-1} is defined as in (23) with the past rejection indicators, the sequence C_1,\ldots,C_{t-1} is defined as in Section 2.1 and $\mathbf{A}_t(\cdot)$ is some functional. This ensures that the sequence of critical values is predictable, that is, α_t is \mathcal{F}_{t-1} -measurable as defined in Section 2.1. In addition, for better readability, we will omit C_1,\ldots,C_{t-1} in the notation of $\mathbf{A}_t(\cdot)$, writing $\mathbf{A}_t(R_1,\ldots,R_{t-1})$ instead of $\mathbf{A}_t(R_1,\ldots,R_{t-1},C_1,\ldots,C_{t-1})$ in Sections 4.3 and 4.4. This should not cause confusion because the recursion is w.r.t. the critical values α_t , which are not used in the sequence C_1,\ldots,C_{t-1} .

4.1. Warming up: LORD procedure and a first greedy reward

While a sufficient condition for online FWER control is $\sum_{t\geq 1} \alpha_t \leq \alpha$ (see the previous section and in particular (21)), the mFDR control is ensured when $\sum_{t\geq 1} \alpha_t \leq \alpha (1\vee R(T))$, as proved in Theorem 2 of Ramdas et al. (2017) (applicable, e.g., under assumptions (2) and (3)). Consequently, for each rejection we earn back some α wealth with which we are allowed to increase α_t ; typically by starting a new online Bonferroni critical value process. This idea is referred to as α -investing in the literature, see Foster and Stine (2008); Aharoni and Rosset (2014); Javanmard and Montanari (2018). This idea leads to the LORD (Levels based On Recent Discovery) procedure (Javanmard and Montanari, 2018), with the improvement given by Ramdas et al. (2017): the critical values are defined recursively in time $T \geq 1$ as

$$\alpha_T^{\text{LORD}} = \mathbf{A}_T^{\text{LORD}}(R_1, \dots, R_{T-1}) := W_0 \gamma_T + (\alpha - W_0) \gamma_{T-\tau_1} + \alpha \sum_{j \ge 2} \gamma_{T-\tau_j},$$
 (25)

where by convention $\gamma_t = 0$ at any time $t \leq 0$ and where γ is an arbitrary spending sequence. Note that the test level at time T splits the initial α -wealth between the cases where R(T) = 0 and R(T) = 1, because the bound is equal to $\alpha(1 \vee R(T)) = \alpha$ in both cases so the first rejection does not provide an extra room for false discoveries. The resulting additional parameter $W_0 \in (0, \alpha)$ balances the initial α -wealth between these two cases to maintain the mFDR control. The procedure $\mathcal{A}^{\text{LORD}} = \{\alpha_t^{\text{LORD}}, t \geq 1\}$ controls the mFDR under (1) and (3), because for any rejection sequence (R_1, \ldots, R_{T-1}) , we have $\sum_{t>1} \mathbf{A}_{\text{L}}^{\text{LORD}}(R_1, \ldots, R_{t-1}) \leq \alpha(1 \vee R(T))$ (see Section 8.2 for a proof).

Now, let us consider our more general framework where we have at hand a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2). In that case, Theorem 4.3 below implies that mFDR is controlled, if we can ensure that $\widehat{\text{FDP}}(T, \mathcal{A}) \leq \alpha$ for all $T \in \{1, 2, ...\}$, where

$$\widehat{\text{FDP}}(T, \mathcal{A}) := \frac{\alpha_T + \sum_{t=1}^{T-1} F_t(\alpha_t)}{1 \vee R(T)}.$$
 (26)

By using the crude bound $F_t(\alpha_t) \leq \alpha_t$, we recover the classical criterion for mFDR control mentioned in the beginning of this section. Similar to the reasoning described in Section 3.1 for FWER control, there are various ways of ensuring that $\widehat{\text{FDP}}(T, \mathcal{A}) \leq \alpha$. A simple solution that takes the LORD critical values as a benchmark is to solve the equation

$$\alpha_T = \sum_{t=1}^{T} \mathbf{A}_t^{\text{LORD}}(R_1, \dots, R_{t-1}) - \sum_{t=1}^{T-1} F_t(\alpha_t), \quad T \ge 1,$$

where we recall that $R_t = \mathbf{1}\{p_t \leq \alpha_t\}, t \leq T-1$. The solution of this equation can be provided by the following recursion: $\alpha_1 = W_0 \gamma_1$ and given $\alpha_1, \ldots, \alpha_{T-1}$, we let

$$\alpha_T = \mathbf{A}_T^{\text{LORD}}(R_1, \dots, R_{T-1}) + \rho_{T-1}, \quad T \ge 1,$$
 (27)

where $\rho_{T-1} = \alpha_{T-1} - F_{T-1}(\alpha_{T-1})$ is the super-uniformity reward (8) at time T-1 (with the convention $\rho_0 = 0$). Since $\rho_t \geq 0$ for all t by (2), this procedure uniformly dominates the procedure $\mathcal{A}^{\text{LORD}}$. Furthermore, depending on the magnitude of the super-uniformity reward, this new procedure is potentially much more powerful.

Example 4.1. Let us provide an example of critical values built from (27): for the rejection sequence $(R_1, R_2, R_3) = (0, 1, 1)$ (so that $\tau_1 = 2, \tau_2 = 3$), we have $\alpha_1 = W_0 \gamma_1$, $\alpha_2 = W_0 \gamma_2 + \alpha_1 - F_1(\alpha_1)$, $\alpha_3 = W_0 \gamma_3 + (\alpha - W_0) \gamma_1 + \alpha_2 - F_2(\alpha_2)$, $\alpha_4 = W_0 \gamma_4 + (\alpha - W_0) \gamma_2 + \alpha \gamma_1 + \alpha_3 - F_3(\alpha_3)$.

Remark 4.1. It is apparent from (27) that the new critical values can be written as $\alpha_T = \mathbf{A}_T(R_1, \dots, R_{T-1})$ for some functional $\mathbf{A}_T(\cdot)$. However, since the super-uniformity reward (8) also depends on the rejection sequence, this functional has a subtle behavior. As a matter of fact, while the functional $\mathbf{A}_T^{LORD}(\cdot)$ is coordinate-wise non decreasing, this is not necessarily the case for the rewarded version $\mathbf{A}_T(\cdot)$, see Proposition 10.1.

4.2. Smoothing out the super-uniformity reward

As discussed for FWER control (see Section 3.2), the preliminary procedure (27) spends immediately at time T all of the super-uniformity reward collected at time T-1. However, it is more advantageous to redistribute this reward over subsequent times $T, T+1, \ldots$, by using a SUR spending sequence $\gamma' = (\gamma'_t)_{t\geq 1}$. This gives rise to the following more general class of online procedures.

Definition 4.1. For a spending sequence γ and a SUR spending sequence γ' , the LORD procedure with super-uniformity reward, denoted by $\mathcal{A}^{\rho LORD} = \{\alpha_t^{\rho LORD}, t \geq 1\}$, is defined by the recursion

$$\alpha_T^{\rho LORD} = \mathbf{A}_T^{LORD}(R_1, \dots, R_{T-1}) + \sum_{t=1}^{T-1} \gamma'_{T-t} \rho_t, \quad T \ge 1,$$
 (28)

where $\mathbf{A}_{T}^{\text{LORD}}$ is given by (25), $R_{t} = \mathbf{1}\{p_{t} \leq \alpha_{t}^{\rho^{\text{LORD}}}\}$ and $\rho_{t} = \alpha_{t}^{\rho^{\text{LORD}}} - F_{t}(\alpha_{t}^{\rho^{\text{LORD}}})$ denotes the super-uniformity reward at time t.

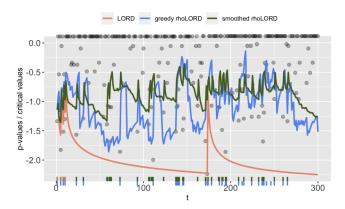


FIG 4. Sequences of critical values of LORD procedure with different rewards over time $1 \le t \le T = 300$ (simulated data): base LORD critical values (25) (orange line), rewarded with the greedy approach (27) (blue line), and with the rectangular kernel SUR spending sequence (28) (h = 10, green line). The rug plots display the time of discoveries for each procedure with the corresponding color. The y-axis has been transformed by $y \mapsto -\log(-\log(y))$. The grey dots denote the p-value sequence (those equal to 1 are displayed at the top of the picture). The spending sequence is $\gamma_t \propto t^{-1.6}$.

Figure 4 displays the critical values of the LORD procedure, and of those rewarded with the greedy SUR spending sequence $\gamma'=(1,0,\dots)$ or rewarded with the rectangular kernel SUR spending sequence (28) (h=10). First, the reward given by the α -investing, which is possible for mFDR control, is visible at each discovery for which all critical value curves 'jump'. Second, the effect of the super-uniformity reward is visible between these jumps, and the kernel sequence is able to better smooth the critical value sequence. As a result, the corresponding procedure is likely to make more discoveries (as in the simulated data presented in Figure 4). The following result establishes the mFDR control of this new class of rewarded procedures.

Theorem 4.1. Consider the setting of Section 2.1 where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. For any spending sequence γ and any SUR spending sequence γ' , consider the LORD procedure $\mathcal{A}^{LORD} = \{\alpha_t^{LORD}, t \geq 1\}$ (25) and the LORD procedure with super-uniformity rewards $\mathcal{A}^{\rho LORD} = \{\alpha_t^{\rho LORD}, t \geq 1\}$ (28). Then, assuming that the model \mathcal{P} is such that (3)

holds, we have $\operatorname{mFDR}(\mathcal{A}^{\rho LORD}, P) \leq \alpha$ for all $P \in \mathcal{P}$ while $\mathcal{A}^{\rho LORD}$ uniformly dominates \mathcal{A}^{LORD} .

This theorem is proved in Section 8.2, as a corollary of a more general result (Theorem 4.3 below). As shown in the numerical experiments (Section 5.2), the improvement of $\mathcal{A}^{\rho \text{LORD}}$ with respect to $\mathcal{A}^{\text{LORD}}$ can be substantial.

Remark 4.2. $A^{\rho LORD}$ can be also expressed by using the paradigm of generalized α investing (GAI) rules, as introduced in Foster and Stine (2008); Aharoni and Rosset (2014); Ramdas et al. (2017), see Section 12.3.

4.3. Rewarded adaptive LORD

In this section, we apply the re-indexation trick of the γ sequence presented in Section 3.3 to improve the performance of the procedures $\mathcal{A}^{\text{LORD}}$ and $\mathcal{A}^{\rho \text{LORD}}$. For this, we follow essentially the reasoning used by Ramdas et al. (2018) for deriving the SAFFRON procedure, with a slight modification, as explained below. To start, let us define, for some parameter $\lambda \in [0, 1)$,

$$\mathcal{T}_j(T) = \begin{cases} 1 + \sum_{t=\tau_j+2}^T \mathbf{1} \{ p_{t-1} \ge \lambda \} & \text{if } T \ge \tau_j + 1 \\ 0 & \text{if } T \le \tau_j \end{cases}, \quad j \ge 1, \quad (29)$$

with $\mathcal{T}_0(T) = \mathcal{T}(T)$ given by (16) by convention. From an intuitive point of view, $\mathcal{T}_j(T)$ is like a 'stopwatch' starting after τ_j and suspended at each time t for which $p_{t-1} < \lambda$. Hence, having $p_t < \lambda$ allows to delay the natural dissipation of α -wealth due to online testing. Then, the SAFFRON procedure (Ramdas et al., 2018) is defined by the threshold

$$\alpha_T = \min\left(\lambda, (1 - \lambda) \left(W_0 \gamma_{\mathcal{T}_0(T)} + (\alpha - W_0) \gamma_{\mathcal{T}_1(T)} + \alpha \sum_{j \ge 2} \gamma_{\mathcal{T}_j(T)}\right)\right). \quad (30)$$

This procedure controls the mFDR and FDR under (1) and (3) as proved by Ramdas et al. (2018). The capping prevents the critical values from exceeding λ , thus avoiding the case $p_t \geq \lambda$ when $p_t \leq \alpha_t$, and makes the procedure monotone in the past decisions, see Lemma 1 in Ramdas et al. (2018). As the monotonicity is not needed for mFDR control, we discard the capping here and work with the (uniformly dominating) procedure

$$\alpha_T^{\text{ALORD}} = \mathbf{A}_T^{\text{ALORD}}(R_1, \dots, R_{T-1})$$

$$= (1 - \lambda) \left(W_0 \gamma_{\mathcal{T}_0(T)} + (\alpha - W_0) \gamma_{\mathcal{T}_1(T)} + \alpha \sum_{j \ge 2} \gamma_{\mathcal{T}_j(T)} \right). \tag{31}$$

The mFDR control of (31) now follows as a special case of Theorem 4.2 below with $F_t(u) = u$ for all t, u. Also note that $\mathcal{A}^{\text{ALORD}}$ reduces to $\mathcal{A}^{\text{LORD}}$ (25) when $\lambda = 0$, because $\mathcal{T}_j(T) = 0 \vee (T - \tau_j)$ in that case. Now, we generalize this method to our present framework.

Definition 4.2. For a spending sequences γ , a SUR spending sequence γ' and $\lambda \in [0,1)$, the adaptive LORD procedure with super-uniformity reward denoted by $\mathcal{A}^{\rho^{ALORD}} = \{\alpha_t^{\rho^{ALORD}}, t \geq 1\}$, is defined by

$$\alpha_T^{\rho_{ALORD}} = \mathbf{A}_T^{ALORD}(R_1, \dots, R_{T-1}) + \sum_{\substack{1 \le t \le T-1 \\ p_t > \lambda}} \gamma_{T-t}' \rho_t + \varepsilon_{T-1}, \quad T \ge 1, \quad (32)$$

where α_T^{ALORD} is defined by (31), $R_t = \mathbf{1}\{p_t \leq \alpha_t^{\rho_{ALORD}}\}$, $\rho_t = \alpha_t^{\rho_{ALORD}} - F_t(\alpha_t^{\rho_{ALORD}})$ denotes the super-uniformity reward a time t and $\varepsilon_{T-1} = \mathbf{1}\{p_{T-1} < \lambda\}(\alpha_{T-1}^{\rho_{ALORD}} - \alpha_{T-1}^{ALORD})$ is an additional 'adaptive' reward at time T-1 (convention $\varepsilon_0 = 0$).

Note that $\mathcal{A}^{\rho\text{ALORD}}$ reduces to $\mathcal{A}^{\rho\text{LORD}}$ (28) when $\lambda = 0$, and to $\mathcal{A}^{\text{ALORD}}$ when $F_t(u) = u$ for all u, t. The following result shows that this class of procedures controls the mFDR.

Theorem 4.2. Consider the setting of Section 2.1 where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. For any spending sequence γ and any SUR spending sequence γ' , consider the adaptive LORD procedure $\mathcal{A}^{ALORD} = \{\alpha_t^{ALORD}, t \geq 1\}$ (31), and the adaptive LORD procedure with super-uniformity rewards $\mathcal{A}^{\rho ALORD} = \{\alpha_t^{\rho ALORD}, t \geq 1\}$ (32). Then, assuming that the model \mathcal{P} is such that (3) holds, we have mFDR($\mathcal{A}^{\rho ALORD}, P$) $\leq \alpha$ for all $P \in \mathcal{P}$ while $\mathcal{A}^{\rho ALORD}$ uniformly dominates \mathcal{A}^{ALORD} and thus also the SAFFRON procedure of Ramdas et al. (2018).

Theorem 4.2 follows from Theorem 4.3 below. Let us underline that $\mathcal{A}^{\rho \text{ALORD}}$ both incorporates α -investing and super-uniformity reward. Thus, it is expected to be the most powerful among the procedures considered in the present paper. This is supported both by the numerical experiments of Section 5.2 and the real data analysis in Section 5.3.

Remark 4.3. Note that the critical values of ALORD and ρ ALORD can exceed 1 (e.g., when all p-values are zero). Since the rejection decision is the same for a critical value larger than 1 or equal to 1, this may appear at first sight as wasted wealth. While this is indeed the case for ALORD, we emphasize that this is not the case for ρ -ALORD, because the super-uniformity reward allows to reuse the exceeding amount of wealth available in $\alpha_t^{\rho_{ALORD}}$; namely $\rho_t = \alpha_t^{\rho_{ALORD}} - 1$ when $\alpha_t^{\rho_{ALORD}} \geq 1$.

4.4. Rewarded version for base mFDR controlling procedures

In this section, we present a result establishing that any base online mFDR controlling procedure satisfying mild conditions (of a specific type) can be rewarded with super-uniformity. First, we can extend (26) by defining

$$\widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) = \frac{\alpha_T + \sum_{\substack{1 \le t \le T-1, \\ p_t \ge \lambda}} F_t(\alpha_t)}{(1 - \lambda)(1 \vee R(T))}.$$
(33)

The following lemma holds (proved in Section 8.2).

Lemma 4.1. Assume that both (2) and (3) hold. For any procedure $\mathcal{A} = (\alpha_t, t \geq 1)$, if for some $\lambda \in [0, 1)$ we have $\sup_{T \geq 1} \widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) \leq \alpha$ a.s. then $\operatorname{mFDR}(\mathcal{A}, P) \leq \alpha$.

This gives rise to the following theorem.

Theorem 4.3. Assume that both (2) and (3) hold. Consider any sequence of functionals $\mathbf{A}_{T}^{0}(\cdot)$, $T \geq 1$, such that for all $R_{1}, \ldots, R_{T-1} \in \{0, 1\}$, we have

$$\mathbf{A}_{T}^{0}(R_{1},\ldots,R_{T-1}) + \sum_{\substack{1 \le t \le T-1, \\ p_{t} \ge \lambda}} \mathbf{A}_{t}^{0}(R_{1},\ldots,R_{t-1}) \le (1-\lambda)\alpha (1 \lor R(T)), \quad (34)$$

where $R(T) = \sum_{t=1}^{T} R_t$, and consider the corresponding procedure $\mathcal{A}^0 = (\alpha_t^0, t \geq 1)$ defined recursively by $\alpha_T^0 = \mathbf{A}_T^0(R_1^0, \dots, R_{T-1}^0)$, $T \geq 1$, with $R_t^0 = \mathbf{1}\{p_t \leq \alpha_t^0\}$, $t \leq T-1$. Then the following holds

- (i) \mathcal{A}^0 controls the online mFDR, that is, mFDR(\mathcal{A}^0, P) $\leq \alpha$ for all $P \in \mathcal{P}$;
- (ii) for any SUR spending sequence $\gamma' = (\gamma'_t, t \ge 1)$, the procedure $\mathcal{A} = (\alpha_t, t \ge 1)$, corresponding to the rewarded \mathcal{A}^0 , and defined by the recursion

$$\alpha_{T} = \mathbf{A}_{T}^{0}(R_{1}, \dots, R_{T-1}) + \sum_{\substack{1 \leq t \leq T-1 \\ p_{t} \geq \lambda}} \gamma'_{T-t}(\alpha_{t} - F_{t}(\alpha_{t})) + \mathbf{1}\{p_{T-1} < \lambda\}(\alpha_{T-1} - \mathbf{A}_{T-1}^{0}(R_{1}, \dots, R_{T-2})), \quad T \geq 1,$$
 (35)

for $R_t = \mathbf{1}\{p_t \leq \alpha_t\}$, controls the online mFDR, that is, mFDR(\mathcal{A}, P) $\leq \alpha$ for all $P \in \mathcal{P}$.

Theorem 4.3 is proved in Section 8.2. Condition (34) is essentially the same as the condition found in Theorem 1 of Ramdas et al. (2018). Our main contribution is in statement (ii), showing that the super-uniformity reward can be used with any base procedure \mathcal{A}^0 satisfying (34). Since the latter condition holds for the LORD procedure $\mathcal{A}^0 = \mathcal{A}^{\text{LORD}}$, and the adaptive LORD procedure $\mathcal{A}^0 = \mathcal{A}^{\text{ALORD}}$ (see Lemma 8.3), Theorem 4.3 entails Theorem 4.1 and Theorem 4.2, respectively. Finally, let us emphasize the similarity between Theorem 3.3 (FWER) and Theorem 4.3 (mFDR). Interestingly, the reward takes exactly the same form (22), which makes the range of improvement in the critical values comparable for these two criteria.

Remark 4.4. The results of Theorem 4.3 is also true for the mFDR taken at some stopping time, see Appendix 9.

Remark 4.5. While this section considers the online mFDR control, we show in Appendix 10 that our SUR approach can also be developed for online (non-marginal) FDR control, see Theorem 10.2 and the modified procedure $\mathcal{A}^{\rho LORD}$ in Definition 10.1. The procedure is developed in the non-adaptive setting. It uniformly improves \mathcal{A}^{LORD} and is dominated by $\mathcal{A}^{\rho LORD}$. However, computing the corresponding reward becomes rapidly intractable in time so the interest in this procedure is mainly theoretical.

5. SUR procedures for discrete tests

In this section, we study the performances of our newly derived SUR procedures in discrete online multiple testing problems for simulated and real data. We defer some of the numerical results to Appendix 14.

5.1. Considered procedures

The considered procedures are the base (non-rewarded) procedures \mathcal{A}^{OB} (10), \mathcal{A}^{AOB} (17), $\mathcal{A}^{\text{LORD}}$ (25), and $\mathcal{A}^{\text{ALORD}}$ (31), and their rewarded counterparts \mathcal{A}^{POB} (18), $\mathcal{A}^{\text{PAOB}}$ (18), $\mathcal{A}^{\text{PLORD}}$ (32), and $\mathcal{A}^{\text{PALORD}}$ (32), respectively. As mentioned in Section 1.5, we also consider the ADDIS-spending and ADDIS procedures (see Tian and Ramdas, 2021, 2019) although the type I error rate control is not guaranteed for these two procedures, in our (discrete) setting. The parameters of the OMT procedures are set to $\alpha = 0.2$, $W_0 = \alpha/2$ and $\lambda = 0.5$. For ADDIS and ADDIS-spending, we use the default values $W_0 = \frac{\alpha \lambda \tau}{2}$, with $\lambda = 0.25$ and $\tau = 0.5$ (the latter being the discarding parameter, see Tian and Ramdas, 2021, 2019). Following Tian and Ramdas (2019), we set $\gamma_t \propto t^{-1.6}$ with a normalizing constant chosen such that $\sum_{t=1}^{+\infty} \gamma_t = 1$. For the SUR spending sequence $(\gamma_t')_{t\geq 1}$ we use a rectangular kernel with bandwidth h, as defined by (9), with h = 100 for FWER and h = 10 for mFDR. We discuss different choices for tuning parameters in the SUR procedures (adaptivity parameter λ and the rectangular kernel bandwidth h) in Appendices 14.4 and 14.5.

5.2. Application to simulated data

5.2.1. Simulation setting

We simulate m experiments in which the goal is to detect differences between two groups by counting the number of successes/failures in each group. More specifically, we follow Gilbert (2005b), Heller and Gur (2011) and Döhler et al. (2018) by simulating a two-sample problem in which a vector of m independent binary responses is observed for N subjects in both groups. The goal is to test the m null hypotheses H_{0i} : ' $p_{1i} = p_{2i}$ ', i = 1, ..., m in an online fashion, where p_{1i} and p_{2i} are the success probabilities for the i^{th} binary response in group A and B respectively. Thus, for each hypothesis i, the data can be summarized by a 2×2 contingency table, and we use (two-sided) Fisher's exact test for testing H_{0i} . The m hypotheses are split in three groups of size m_1, m_2 , and m_3 such that $m = m_1 + m_2 + m_3$. Then, the binary responses are generated as i.i.d Bernoulli of probability 0.01 ($\mathcal{B}(0.01)$) at m_1 positions for both groups, i.i.d $\mathcal{B}(0.10)$ at m_2 positions for both groups, and i.i.d $\mathcal{B}(0.10)$ at m_3 positions for one group and i.i.d $\mathcal{B}(p_3)$ at m_3 positions for the other group. Thus, the null hypotheses are true for $m_1 + m_2$ positions (set \mathcal{H}_0), while the null hypotheses are false for m_3 positions (set \mathcal{H}_1). Therefore, we interpret p_3 as the strength of the signal while $\pi_A = \frac{m_3}{m}$, corresponds to the proportion of signal. Also, m_1 and m_2 are both

taken equal to $\frac{m-m_3}{2}$. In these experiments, we fix m=500, and vary each one of the parameters \mathcal{H}_1 (Section 5.2.2), π_A (Section 5.2.3), N (Section 14.1), p_3 (Section 14.2) while keeping the others fixed. The default values are $\pi_A=0.3$, N=25, $p_3=0.4$ and $\mathcal{H}_1 \subset \{1,\ldots,m\}$ chosen randomly for each simulation run. We estimate the different criteria (FWER (5), mFDR (6), power (7)) using empirical mean over 10 000 independent simulation trials.

5.2.2. Position of signal

We start by studying how the position of the signal can affect the performances of the procedures (it is well-known to be critical, see Foster and Stine, 2008; Ramdas et al., 2017). We investigate different positioning schemes in which the signal can be clustered at the beginning of the stream, or at the end, or clustered between the two, as described in the caption of Figure 5.

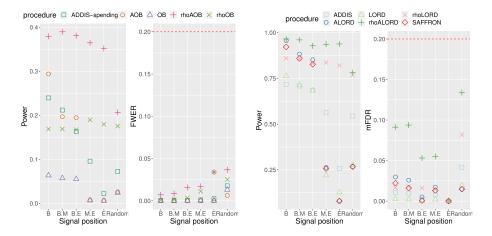


FIG 5. Power and type I error rates of the different considered OMT procedures versus positions of the signal: at the beginning (B), the end (E), half at the beginning and half in the middle of the stream (BM), half at the beginning and half at the end of the stream (BE), half in the middle and half at the end of the stream (ME), and taken uniformly at random (Random).

Consistently with our theoretical results, Figure 5 shows that all procedures control the type I error rate at level $\alpha=0.2$. In terms of power, we can see that the rewarded procedures have greater power than the associated base procedures. More specifically, $\mathcal{A}^{\rho\text{ALORD}}$ uniformly dominates the other procedures for mFDR control and $\mathcal{A}^{\rho\text{AOB}}$ for FWER control. The gain in power is most noticeable when the signal is not localized at the beginning of the stream (i.e. positions ME, E, and Random) for which the online testing problem is more difficult. These first results indicate that the rewarded procedures may protect against ' α -death'.

5.2.3. Proportion of signal

Figure 6 displays the results for π_A varying in $\{0.1, \ldots, 1\}$. It shows that the aforementioned superiority of the rewarded procedures holds in this whole range. Also note that the super-uniformity reward can affect the monotonicity of the power curves: while most curves are increasing with π_A , the power of the rewarded procedure $\mathcal{A}^{\rho \text{OB}}$ decreases. An explanation could be that when π_A increases, the marginal counts increase, and thus the degree of discreteness decreases providing a smaller super-uniformity reward. However, using adaptivity seems to compensate for this effect, thus providing better results.

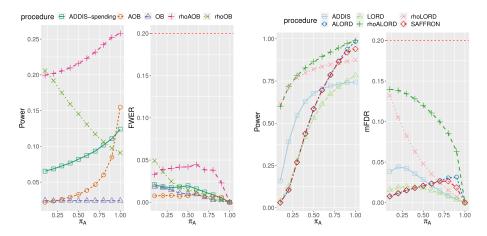


Fig 6. Power and type I error rates of the considered procedures for $\pi_A \in \{0.1, 0.2, \dots, 0.9, 1\}$.

Finally, let us mention that the additional numerical results in Section 14 provide qualitatively similar conclusions for all other explored parameter configurations: the SUR procedures $\mathcal{A}^{\rho \text{AOB}}$ and $\mathcal{A}^{\rho \text{ALORD}}$ always improve, often substantially, the existing OMT procedures.

5.3. Application to IMPC data

In this section we analyse data from the International Mouse Phenotyping Consortium (IMPC), which coordinates studies on the genotype influence on mouse phenotype. More precisely, scientists test the hypotheses that the knock-out of certain genes will not change certain phenotypic traits (e.g., the coat or eye color). Since the data set is constantly evolving as new genes are studied for new phenotypic traits of interest, online multiple testing is a natural approach for analysing such data, see also Tian and Ramdas (2021); Xu and Ramdas (2021). We use the data set provided by Karp et al. (2017) which includes, for each studied gene, the count of normal and abnormal phenotype for female and male mice (separately), thus providing two by two contingency tables, which can be analysed using Fisher exact tests. In this section, we investigate the

genotype effect on the phenotype separately for male and female. The data set originally contains nearly 270 000 genes studies, but we focus on the first 30 000 genes for simplicity. We set the global level α to 0.2 and 0.05, respectively for FWER and mFDR procedures. For the procedure parameters, we follow the choice made in Section 5.1. Table 3 presents the number of discoveries for the FWER controlling procedures OB, AOB, ρ OB, ρ AOB (top) and for the mFDR controlling procedures LORD, ALORD, ρ LORD, ρ ALORD (bottom). The results show that ignoring the discreteness of the tests causes the scientist to miss (potentially many) discoveries. Hence, using the SUR methods helps to reduce this risk.

Table 3
Number of discoveries for FWER controlling OMT procedures (top) and mFDR controlling OMT procedures (bottom). These numbers are obtained by running the procedures on the first 30 000 genes for male (second row) and female (third row) mice in the IMPC data.

FWER procedures	OB	$ ho \mathrm{OB}$	AOB	ho AOB
# discoveries (male)	229	377	281	697
# discoveries (female)	267	481	764	811
mFDR procedures	LORD	ρ LORD	ALORD	ρ ALORD
# discoveries (male)	882	972	972	1041
# discoveries (female)	839	946	966	1046

Figure 7 (FWER procedures) and Figure 8 (mFDR procedures) illustrate in more detail how the super-uniformity reward leads to more discoveries, in the case of male mice (similar findings hold for the female mice for which the corresponding figures can be found in Section 15.2). First, note that the smallest p-values occur at the beginning of the stream (see Figure 17 in Section 15.1), so that we limit the visual analysis to the first 1500 p-values for clarity of exposition. For the ρ OB procedure, the benefit of incorporating the super-uniformity reward is visible in the left panel of Figure 7. As expected from Figure 3, applying a rectangular kernel to these rewards yields a smooth curve. For the ρAOB procedure, presented in the right panel of Figure 7, the improvement is even stronger, but the resulting critical value curve is less smooth. This is due to the 'adaptive' reward, that is, the ε_{T-1} -component of our improvement, recall (18). More precisely, an explanation of this 'saw-tooth' shape is that during a period with p-values smaller than λ , we have $\alpha_T^{\rho \text{AOB}} - \alpha_T^{\text{AOB}} \geq \alpha_{T-1}^{\rho \text{AOB}} - \alpha_{T-1}^{\text{AOB}}$ so the gain increases. Also, if this period lasts for a while (as for $500 \lesssim t \lesssim 1240$ here), the ρ -part of the reward vanishes and we end up with a constant gain $\alpha_T^{\rho AOB} - \alpha_T^{AOB} \approx \alpha_{T-1}^{\rho AOB} - \alpha_{T-1}^{AOB}$, explaining the flat part of the curve, until the next $p_T \geq \lambda$ occurs. After this point, we switch from the ε -regime back to the ρ -regime, i.e., $\alpha_{T+1}^{\rho \text{AOB}} = \alpha_{T+1}^{\text{AOB}} + \gamma_1' \rho_T$. Since typically $\gamma_1' \rho_T \ll \alpha_{T-1}^{\rho \text{AOB}} - \alpha_{T-1}^{\text{AOB}}$, this causes the downward jump in the green curve. For the mFDR procedures presented in Figure 8, there is an additional 'rejection' reward as described in Section 4. Note that this makes some critical values exceed 1 (both for ALORD and ρ ALORD), which thus cannot be displayed in the Y-axis scale considered in that figure. However, these values are still used in ρ ALORD algorithm to

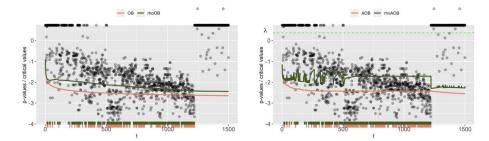


FIG 7. Applying online FWER controlling procedures to the male mice IMPC data set. Left panel: p-values and critical values for OB (orange curve) and ρ OB (green curve). Right panel: AOB (orange curve) and ρ AOB (green curve). Representation similar to Figure 3 (Y-axis transformed by $y \mapsto -\log(-\log(y))$; p-values equal to 1 displayed at the top of the picture).

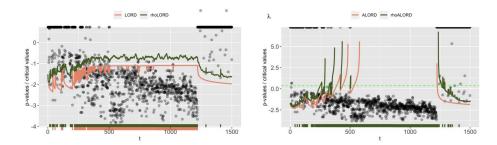


FIG 8. Applying online mFDR controlling procedures to the male mice IMPC data set. Left panel: p-values and critical values for LORD (orange curve) and ρ LORD (green curve). Right panel: ALORD (orange curve) and ρ ALORD (green curve). Representation similar to Figure 4 (Y-axis transformed by $y \mapsto -\log(-\log(y))$; p-values equal to 1 are displayed at the top of the picture).

compute the future critical values (see Remark 4.3). The obtained results are qualitatively similar to the FWER setting: the super-uniformity reward makes the green curves run above the orange ones, uniformly over the considered time, hence inducing significantly more discoveries.

5.4. Discussion

In Sections 3 and 4 we proved that SUR methods are always more powerful than the classical base procedures and the results in this section show that the gains in power can be considerable for discrete tests. Thus, using SUR methods whenever discreteness is involved in tests is a risk-less strategy with potentially large benefits. A more complicated task is to quantify the increase in power that may be expected from using SUR in such a setting. While an exhaustive solution to this problem seems beyond the scope of this paper, we would like to point out some sufficient conditions for when we anticipate SUR to be particularly beneficial:

- When tests with relatively small sample sizes are expected to occur throughout the experiment. The IMPC data can be seen as a typical example of such an experiment, since the number of knockout mice available in each test may be restricted by ethical and financial considerations.
- When discrete tests are involved at the beginning of the OMT process, particularly when using the OB procedure. In this case, the critical value sequence decreases quickly in the beginning and unused wealth can not be recovered later.
- Since we expect many critical values to be small, we may expect SUR to be particularly helpful if the deviations of the F_i 's to the uniform distribution occur for small p-values. The numerical experiments for 2×2 tables in Appendix 14.3 show that even for large sample sizes, where standard rules of thumb would suggest that FET p-values are uniform, the SUR approach still achieves noticeably more power.

6. SUR procedures for weighted p-values

In this section, we show how our SUR approach can be easily used to construct valid online p-value weighting schemes.

6.1. Setting and benchmark procedure

Consider a standard continuous online multiple testing setting where each p-value is super-uniformly distributed under the null, i.e. (1) holds. Assume in addition that, at each time t, the p-value p_t is associated with a quantity $r_t \geq 0$, called the raw weight (as opposed to the rescaled weight defined further on), which is assumed to be measurable w.r.t. \mathcal{F}_{t-1} . Throughout the section, the weights r_t are assumed to be available a priori and we will not discuss how to derive them (for this task, we refer to Wasserman and Roeder (2006); Rubin et al. (2006); Roquain and van de Wiel (2009); Hu et al. (2010); Zhao and Zhang (2014); Ignatiadis et al. (2016); Chen and Kasiviswanathan (2020b) among others).

While p-value weighting is a classical tool for improving the performance of multiple testing methods in the offline setting (see references in Section 1.3), the incorporation of weights has received little attention in the online context. The only relevant work to our knowledge is Ramdas et al. (2017) (Section 5 therein), which presents sufficient criteria for weighting procedures controlling the (m)FDR based on so-called GAI++ procedures and also discusses the technical challenges associated with weighted online multiple testing procedures. An explicit algorithm which satisfies these criteria is used in Ramdas et al. (2017)⁴, which is detailed in Appendix 12.2 for completeness. This method, which will be our benchmark procedure, works by weighting the p-values and adjusting for this weighting in the rejection reward.

⁴An implementation of this procedure can be found on the website https://github.com/fanny-yang/OnlineFDRCode

6.2. New weighting approach

The main idea of our new approach is as follows: consider weighted p-values $\tilde{p}_t = p_t/w_t$ for some rescaled weight $w_t \in [0,1]$ which gives rise to the null bounding family $\mathfrak{F} = \{F_t : u \in [0,1] \mapsto uw_t, t \geq 1\}$. Since the weights are constrained to take their values in [0,1], the functions of \mathfrak{F} are super-uniform, i.e. (2) holds. Hence, one can apply our SUR approach with respect to that family \mathfrak{F} .

More specifically, our approach takes into account the null bounding family \mathfrak{F} in a simple two-step process, which proceeds as follows: for each time t,

1. enforce super-uniformity by computing the rescaled weight

$$w_t = \xi_t(r_t|r_1, \dots, r_{t-1}),$$

- $t \geq 1$, for some given rescaling function ξ_t valued in [0,1] (see below for more details and an explicit choice);
- 2. apply any one of the SUR methods from Section 3 or Section 4, depending on whether FWER or mFDR control is desired.

The second step of the above algorithm is equivalent to rejecting hypothesis H_t if $p_t \leq w_t \alpha_t$, so that w_t is the fraction of α_t spent on testing H_t and $1 - w_t$ is the fraction of α_t saved for later testing. In this view, it is appropriate to interpret the quantity $1 - w_t$ in our approach as a penalty weight at time t, which serves to make a rejection of H_t less likely. This is different from conventional offline weighting where the original p-values can be directly up-weighted or down-weighted. Here, the online weighting acts in a more subtle way in the decision rule $p_t \leq w_t \alpha_t$; while $w_t \alpha_t$ is down-weighted by w_t (because $w_t \in [0,1]$), it is up-weighted by past weights w_1, \ldots, w_{t-1} .

At first sight, these SUR weighting approaches may seem to be ineffective due to the conservatism induced by the rescaling step. However, this is countered in the second step by using SUR procedures that provide larger values α_t , due to the super-uniform rewards accumulated in the past. The hope is that these two effects balance out in such a way as to favor rejection of hypotheses associated with larger values of (raw) weights.

We denote these new weighted procedures by wX, where X stands for the name of the base procedure (either OB (10), AOB (17), LORD (25) or ALORD (31)). These procedures all come with the corresponding FWER or mFDR control (by additionally assuming (3) if needed). In particular, to the best of our knowledge, this also provides the first method for weighted online FWER control.

Finally, let us mention that a simple choice for ξ_t is given by $\xi_t(x|r_1,\ldots,r_{t-1}) = \hat{F}_{t-1}(x)\mathbf{1}\{x>0\}$, where $\hat{F}_{t-1}(x) = (t-1)^{-1}\sum_{i=1}^{t-1}\mathbf{1}\{r_i \leq x\}$ is the empirical c.d.f. of the sample r_1,\ldots,r_{t-1} (and by convention $\hat{F}_0(x)=1$). This particular choice is easy to compute in a sequential manner, and it satisfies the following intuitive and desirable properties: $\xi_t(x) \in [0,1]$ (ensures super-uniformity of \mathfrak{F}), $\xi_t(x)$ is nondecreasing in x (a larger raw weight leads to a larger rescaled

weight), $\xi_t(0) = 0$ (raw zero weights rescaled to zero), $\xi_t(\lambda r_t | \lambda r_1, \dots, \lambda r_{t-1}) = \xi_t(r_t | r_1, \dots, r_{t-1})$ for all $\lambda > 0$ (scale invariance) and if all raw weights are equal then all rescaled weights are equal to 1.

6.3. Analysis of RNA-Seq data

We revisit an analysis of the RNA-Seq data set 'airway' using results from the Independent Hypothesis Weighting (IHW) approach (for details, see Ignatiadis et al. (2016) and the vignette accompanying its software implementation). While the original data was not collected in an online fashion, we use it here nevertheless to provide a proof of concept for weighted SUR procedures. The 'airway' data set contains data from 64102 genes and the corresponding (offline) weights are taken from the output of the ihw function from the bioconductor package 'IHW'. These 'raw' weights are then transformed into rescaled weights by using the function ξ_t described in the previous section. For the procedure parameters, we use the same choices as for the analysis of the IMPC data, see Section 5.3.

Table 4 (top part) presents the result for the FWER controlling procedures OB, AOB (non-weighted), and wOB, wAOB (SUR weighted approaches). It is clear that incorporating the weights leads to more rejections, which corroborates the fact that the weights coming from Ignatiadis et al. (2016) are indeed informative.

Table 4
Number of discoveries for weighted controlling OMT procedures for the 'airway' data set, with the weights taken from Ignatiadis et al. (2016).

FWER procedures	OB	wOB (new)	AOB	wAOB (new)
# discoveries	1092	1195	1188	1273
mFDR procedures	LORD	$w\mathrm{GAI}_1$	$wGAI_2$	wLORD (new)
# discoveries	3550	1308	3631	3875

As for mFDR control, the (non-weighted) LORD is compared to our weighted version wLORD in Table 4 (bottom part). As additional competitors, we also added the weighted GAI++ procedure proposed in Ramdas et al. (2017) (see Section 12.2 for a detailed description), that we use either with the raw weights (denoted by wGAI₁) or with the rescaled weights (denoted by wGAI₂). As one can see, the effect of rescaling the weights is highly beneficial, and the new wLORD proposal is the one that incorporates these weights in the most efficient way.

7. Discussion

7.1. Conclusion

Existing OMT procedures often suffer from a lack of power due to conservativeness of the p-values. This occurs typically for discrete test statistics, which is a common situation when testing is based upon counts. To fill the gap, we

introduced new SUR versions of some existing classical procedures, that 'reward' the base procedures by spending more efficiently the α -wealth according to known bounds on the null cumulative distribution functions. We showed that our new SUR procedures provide rigorous control of online error criteria (FWER or mFDR) under classical assumptions while offering a systematic power enhancement. When using discrete Fisher exact test statistics, the improvement is substantial, both for simulated and real data.

In addition, even in the standard case of uniformly distributed *p*-values, our approach allowed us to derive new weighted procedures that incorporate external covariates. This provides improvements w.r.t. existing online weighting strategies.

7.2. Future directions

While our results address several issues, they also raise new questions. First, the bandwidth of the kernel-based SUR spending sequence γ' given by (9) has been chosen in a loose way here, but tuning the bandwidth is certainly interesting from a power enhancement perspective (see Section 14.5). Also, in applications, the user would possibly like to select the bandwidth in a data dependent fashion without losing control over type I error rate. These two issues are interesting extensions for future developments. Second, most of our results rely on an independence assumption, see (3). While this can be considered as a mild restriction in an online framework, relaxing it or incorporating a known dependence structure in OMT is an interesting avenue for future work.

8. Proofs

8.1. Proofs for online FWER control

We start by proving Theorem 3.3 and then deduce Theorems 3.1 and 3.2.

Proof of Theorem 3.3. From Lemma 3.1, (i) is clear by using the (crude) bound $F_t(x) \leq x$ and assumption (21). As for (ii), by using again Lemma 3.1, we only have to check that the procedure \mathcal{A} defined by (22) satisfies $\widehat{\text{FWER}}_{\lambda}(T,\mathcal{A}) \leq \alpha$ (a.s.). This is done by reducing this to a statement on α_t^0 via Lemma 8.2. More precisely, with $a_T = \sum_{t=1}^T \gamma_t'$, we have

$$\alpha_{T} + \sum_{t=1}^{T-1} \mathbf{1} \{ p_{t}(X) \ge \lambda \} F_{t}(\alpha_{t}) \le \alpha_{T} + \sum_{t=1}^{T-1} \mathbf{1} \{ p_{t} \ge \lambda \} \left[(1 - a_{T-t}) \alpha_{t} + a_{T-t} F_{t}(\alpha_{t}) \right]$$
$$= \alpha_{T}^{0} + \sum_{t=1}^{T-1} \mathbf{1} \{ p_{t} \ge \lambda \} \alpha_{t}^{0} \le \alpha,$$

where the equality above is true provided that the following recursion holds for all $T \ge 1$,

$$\alpha_T = \alpha_T^0 + \sum_{t=1}^{T-1} \mathbf{1} \{ p_t \ge \lambda \} \alpha_t^0 - \sum_{t=1}^{T-1} \mathbf{1} \{ p_t \ge \lambda \} \left[(1 - a_{T-t}) \alpha_t + a_{T-t} F_t(\alpha_t) \right].$$

This is true by Lemma 8.2 because of the expression (22) of α_t . This concludes the proof.

Proof of Theorems 3.1 and 3.2. Theorems 3.1 and 3.2 are corollaries of Theorem 3.3, by considering $\mathcal{A}^0 = \mathcal{A}^{\text{OB}}$ ($\lambda = 0$) and $\mathcal{A}^0 = \mathcal{A}^{\text{AOB}}$, respectively. Indeed, checking (21) is straightforward for \mathcal{A}^{OB} from the spending sequence definition or comes from Lemma 8.3 for \mathcal{A}^{AOB} .

8.2. Proofs for online mFDR control

The global proof strategy is similar to the one used for FWER: we start by proving Theorem 4.3 and then deduce Theorem 4.1 and Theorem 4.2, and finally prove Lemma 4.1.

Proof of Theorem 4.3. By Lemma 4.1 it is enough to prove $\sup_{T\geq 1} \widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) \leq \alpha$ (a.s.) both for $\mathcal{A} = \mathcal{A}^0$ and \mathcal{A} being the rewarded procedure (35). First, it is clear for \mathcal{A}^0 by (34). Second, we can show that condition $\sup_{T\geq 1} \widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) \leq \alpha$ (a.s.) holds for any procedure $\mathcal{A} = (\alpha_t, t \geq 1)$ such that for all $T \geq 1$,

$$\alpha_T + \sum_{t=1}^{T-1} \mathbf{1} \{ p_t \ge \lambda \} \left[(1 - a_{T-t}) \alpha_t + a_{T-t} F_t(\alpha_t) \right]$$

$$= \mathbf{A}_T^0(R_1, \dots, R_{T-1}) + \sum_{p_t > \lambda, 1 \le t \le T-1} \mathbf{A}_{t-1}^0(R_1, \dots, R_{t-1}), \tag{36}$$

where $a_T = \sum_{t=1}^T \gamma_t'$ and $R_t = \mathbf{1}\{p_t \leq \alpha_t\}$. Indeed, we have $\mathbf{A}_T^0(R_1, \dots, R_{T-1}) + \sum_{p_t \geq \lambda, 1 \leq t \leq T-1} \mathbf{A}_{t-1}^0(R_1, \dots, R_{t-1}) \leq (1-\lambda)\alpha(1 \vee R(T))$ by (34), which means $\sup_{T \geq 1} \widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) \leq \alpha$. Now, (36) holds true for the rewarded procedure (35) by Lemma 8.2 (applied⁵ with $\alpha_T^0 = \mathbf{A}_T^0(R_1, \dots, R_{T-1})$), which concludes the proof.

Proof of Theorems 4.1 and 4.2. Theorem 4.1 and Theorem 4.2 can be derived from Theorem 4.3 for $\mathbf{A}_T^0 = \mathbf{A}_T^{\text{LORD}}$ (25) (using $\lambda = 0$) and $\mathbf{A}_T^0 = \mathbf{A}_T^{\text{ALORD}}$ (31), respectively, by checking (34) in both cases. First, for \mathbf{A}^{LORD} , we have for all $R_1, \ldots, R_{T-1} \in \{0, 1\}$,

$$\sum_{t=1}^{T} \mathbf{A}_{t}^{\text{LORD}}(R_{1}, \dots, R_{t-1}) = \sum_{t=1}^{T} \left(W_{0} \gamma_{t} + (\alpha - W_{0}) \gamma_{t-\tau_{1}} + \alpha \sum_{j \geq 2} \gamma_{t-\tau_{j}} \right)$$

⁵Note that these critical values are slightly different from those of \mathcal{A}^0 because the definition of R_t is different from R_t^0 .

$$= W_0 \sum_{t=1}^{T} \gamma_t + (\alpha - W_0) \sum_{t=1}^{T} \gamma_{t-\tau_1}$$

$$+ \alpha \sum_{j \ge 2} \mathbf{1} \{ T - \tau_j \ge 1 \} \sum_{t=1}^{T} \gamma_{t-\tau_j}$$

$$\le \alpha (1 + 0 \lor (R(T - 1) - 1)) \le \alpha (1 \lor R(T)), \quad (37)$$

because $\tau_j \leq T-1$ is equivalent to $R(T-1) \geq j$ by definition. Second, for $\mathcal{A}^{\text{ALORD}}$, we proceed similarly with the help of Lemma 8.3: by definition (31), we have for all $R_1, \ldots, R_{T-1} \in \{0, 1\}$,

$$(1 - \lambda)^{-1} \left(\mathbf{A}_{T}^{\text{ALORD}}(R_{1}, \dots, R_{T-1}) + \sum_{\substack{1 \leq t \leq T-1, \\ p_{t} \geq \lambda}} \mathbf{A}_{t}^{\text{ALORD}}(R_{1}, \dots, R_{t-1}) \right)$$

$$= W_{0} \left(\gamma_{\mathcal{T}_{0}(T)} + \sum_{\substack{1 \leq t \leq T-1, \\ p_{t} \geq \lambda}} \gamma_{\mathcal{T}_{0}(t)} \right) + (\alpha - W_{0}) \left(\gamma_{\mathcal{T}_{1}(T)} + \sum_{\substack{1 \leq t \leq T-1, \\ p_{t} \geq \lambda}} \gamma_{\mathcal{T}_{1}(t)} \right)$$

$$+ \alpha \sum_{j \geq 2} \mathbf{1} \{ T \geq \tau_{j} + 1 \} \left(\gamma_{\mathcal{T}_{j}(T)} + \sum_{\substack{1 \leq t \leq T-1, \\ p_{t} \geq \lambda}} \gamma_{\mathcal{T}_{j}(t)} \right).$$

Finally, by using (41) and (42), the latter is equal to

$$W_0 \sum_{t=1}^{T_0(T)} \gamma_t + (\alpha - W_0) \sum_{t=1}^{T_1(T)} \gamma_t + \alpha \sum_{j \ge 2} \mathbf{1} \{ T \ge \tau_j + 1 \} \sum_{t=1}^{T_j(T)} \gamma_t$$

$$\le W_0 + \alpha - W_0 + \alpha \sum_{j \ge 2} \mathbf{1} \{ T \ge \tau_j + 1 \} = \alpha (1 + 0 \vee (R(T - 1) - 1))$$

$$\le \alpha (1 \vee R(T)),$$

because $T \ge \tau_j + 1$ if and only if $R(T - 1) \ge j$.

Proof of Lemma 4.1. Let us assume (2), (3) and for all $T \ge 1$, $\widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) \le \alpha$ a.s., that is,

$$\alpha_T + \sum_{\substack{1 \le t \le T-1, \\ p_t \ge \lambda}} F_t(\alpha_t) \le (1 - \lambda)\alpha (1 \lor R(T)), \quad \text{(a.s.)}.$$
 (38)

Now applying Lemma 8.1, we have

$$\mathbf{E}_{X \sim P} \left(\sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_0, p_t \le \alpha_t \} \right) \le (1 - \lambda)^{-1} \mathbf{E} \left(\sum_{t=1}^{T} \mathbf{1} \{ p_t \ge \lambda \} F_t(\alpha_t) \right)$$

$$\le \alpha \mathbf{E} (1 \vee R(T)),$$

by using (38), which is exactly the desired mFDR control.

8.3. Auxiliary lemmas

The following lemma provides a tool for controlling both online FWER and mFDR.

Lemma 8.1. For any procedure $A = (\alpha_t, t \ge 1)$, we have for all $\lambda \in [0, 1)$,

$$\mathbf{E}_{X \sim P} \left(\sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_0, p_t \le \alpha_t \} \right) \le (1 - \lambda)^{-1} \mathbf{E} \left(\sum_{t=1}^{T} \mathbf{1} \{ p_t(X) \ge \lambda \} F_t(\alpha_t) \right), \tag{39}$$

provided that (2) holds and if either (3) holds or if the critical values $(\alpha_t, t \ge 1)$ are deterministic.

Proof. Recall α_t is either deterministic or \mathcal{F}_{t-1} -measurable (in which case it is independent of $p_t(X)$ under (3)). Therefore, under the conditions of the lemma, we have in any case: for all $t \in \mathcal{H}_0$, both

$$\mathbf{E}\left(\frac{\mathbf{1}\{p_t(X) > \lambda\}}{1 - \lambda} \middle| \alpha_t\right) \ge 1, \quad \mathbf{P}\left(p_t(X) \le \alpha_t \mid \alpha_t\right) \le F_t(\alpha_t).$$

This entails

$$\mathbf{E}_{X \sim P} \Big(\sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_{0}, p_{t} \leq \alpha_{t} \} \Big)$$

$$= \sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_{0} \} \mathbf{E} (\mathbf{P}(p_{t}(X) \leq \alpha_{t} \mid \alpha_{t}))$$

$$\leq \sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_{0} \} \mathbf{E} (F_{t}(\alpha_{t}))$$

$$\leq \sum_{t=1}^{T} \mathbf{1} \{ t \in \mathcal{H}_{0} \} \mathbf{E} \left(F_{t}(\alpha_{t}) \mathbf{E} \left(\frac{\mathbf{1} \{ p_{t}(X) \geq \lambda \}}{1 - \lambda} \mid \alpha_{t} \right) \right)$$

$$\leq (1 - \lambda)^{-1} \mathbf{E} \left(\sum_{t=1}^{T} \mathbf{1} \{ p_{t}(X) \geq \lambda \} F_{t}(\alpha_{t}) \right).$$

The following representation lemma is the key tool for building the new rewarded critical values.

Lemma 8.2. Let $(\alpha_t^0, t \ge 1)$ be any nonnegative sequence. Let $(\tilde{\alpha}_t, t \ge 1)$ be the sequence defined by the recursive relation: for all $T \ge 1$,

$$\tilde{\alpha}_T = \alpha_T^0 + \sum_{t=1}^{T-1} \mathbf{1} \{ p_t \ge \lambda \} \alpha_t^0 - \sum_{t=1}^{T-1} \mathbf{1} \{ p_t \ge \lambda \} \left[(1 - a_{T-t}) \tilde{\alpha}_t + a_{T-t} F_t(\tilde{\alpha}_t) \right], \tag{40}$$

where $a_T = \sum_{t=1}^T \gamma_t'$, $T \ge 1$ for any real values γ_t' , p_t , λ and functions F_t . Let $(\bar{\alpha}_t, t \ge 1)$ be the sequence defined by the recursive relation

$$\bar{\alpha}_T = \alpha_T^0 + \sum_{\substack{1 \le t \le T-1 \\ p_t > \lambda}} \gamma_{T-t}'(\bar{\alpha}_t - F_t(\bar{\alpha}_t)) + \mathbf{1}\{p_{T-1} < \lambda\}(\bar{\alpha}_{T-1} - \alpha_{T-1}^0), \quad T \ge 1.$$

Then we have $\tilde{\alpha}_t = \bar{\alpha}_t$ for all $t \geq 1$. Moreover, $\bar{\alpha}_t \geq \alpha_t^0$ for all $t \geq 1$ under (2). In particular, these critical values are nonnegative.

Proof. Clearly, $\tilde{\alpha}_1 = \alpha_1^0 = \bar{\alpha}_1$ so the result is satisfied for T = 1. For $T \geq 2$, by using (40) for $\tilde{\alpha}_T$ and $\tilde{\alpha}_{T-1}$, we have

$$\tilde{\alpha}_{T} - \tilde{\alpha}_{T-1} = \alpha_{T}^{0} - \alpha_{T-1}^{0} + \mathbf{1} \{ p_{T-1} \ge \lambda \} \alpha_{T-1}^{0}$$

$$- \mathbf{1} \{ p_{T-1} \ge \lambda \} \left[(1 - a_{1}) \tilde{\alpha}_{T-1} + a_{1} F_{T-1} (\tilde{\alpha}_{T-1}) \right]$$

$$+ \sum_{t=1}^{T-2} \mathbf{1} \{ p_{t} \ge \lambda \} \left[(a_{T-t} - a_{T-t-1}) \tilde{\alpha}_{t} - (a_{T-t} - a_{T-t-1}) F_{t} (\tilde{\alpha}_{t}) \right].$$

Hence, by using $\tilde{\alpha}_{T-1} = \tilde{\alpha}_{T-1} \mathbf{1} \{ p_{T-1} < \lambda \} + \tilde{\alpha}_{T-1} \mathbf{1} \{ p_{T-1} \ge \lambda \}$, we obtain

$$\tilde{\alpha}_{T} = \alpha_{T}^{0} - \mathbf{1} \{ p_{T-1} < \lambda \} \alpha_{T-1}^{0} + \tilde{\alpha}_{T-1} \mathbf{1} \{ p_{T-1} < \lambda \}$$

$$+ \mathbf{1} \{ p_{T-1} \ge \lambda \} \left[\gamma_{1}' \tilde{\alpha}_{T-1} - \gamma_{1}' F_{T-1} (\tilde{\alpha}_{T-1}) \right]$$

$$+ \sum_{t=1}^{T-2} \mathbf{1} \{ p_{t} \ge \lambda \} \left[\gamma_{T-t}' \tilde{\alpha}_{t} - \gamma_{T-t}' F_{t} (\tilde{\alpha}_{t}) \right],$$

because $\gamma'_1 = a_1$, and we recognize the expression given in the lemma.

Let us finally prove that $\bar{\alpha}_T \geq \bar{\alpha}_T^0$ for all $T \geq 1$. This is true for $\bar{\alpha}_1$ because $\bar{\alpha}_1 = \alpha_1^0$. Now, if $\bar{\alpha}_1 \geq \alpha_1^0, \ldots, \bar{\alpha}_{T-1} \geq \alpha_{T-1}^0$ then we also have

$$\bar{\alpha}_T = \alpha_T^0 + \sum_{\substack{1 \le t \le T-1 \\ p_t \ge \lambda}} \gamma'_{T-t} (\bar{\alpha}_t - F_t(\bar{\alpha}_t)) + \mathbf{1} \{ p_{T-1} < \lambda \} (\bar{\alpha}_{T-1} - \alpha_{T-1}^0) \ge \alpha_T^0,$$

because $\bar{\alpha}_t \geq F_t(\bar{\alpha}_t)$ by (2). This finishes the proof.

We now establish a result for the functionals $\mathcal{T}(\cdot)$ and $\mathcal{T}_j(\cdot)$, $j \geq 1$, which are used by the adaptive procedures \mathcal{A}^{AOB} and $\mathcal{A}^{\text{ALORD}}$, respectively.

Lemma 8.3. Consider the functional $\mathcal{T}(\cdot)$ defined by (16) for some realization of the p-values and some $\lambda \in [0,1)$. Then for any sequence $(\gamma_t)_{t\geq 1}$ and for any $T\geq 1$, we have

$$\sum_{t=1}^{T} \mathbf{1} \{ p_t \ge \lambda \} \gamma_{\mathcal{T}(t)} = \sum_{t=1}^{\mathcal{T}(T+1)-1} \gamma_t.$$
 (41)

In addition, for any $j \geq 1$, consider the τ_j defined by (24) and the functional $\mathcal{T}_j(\cdot)$ defined by (29). Then for all $T \geq \tau_j + 1$,

$$\sum_{\substack{1 \le t \le T \\ p_t > \lambda}} \gamma_{\mathcal{T}_j(t)} = \sum_{t=1}^{\mathcal{T}_j(T+1)-1} \gamma_t. \tag{42}$$

Proof. Let us first prove (41). Since $\mathcal{T}(t+1) = \mathcal{T}(t) + 1$ when $p_t \geq \lambda$ from definition (16), we can write

$$\sum_{t=1}^{T} \mathbf{1} \{ p_t \ge \lambda \} \gamma_{\mathcal{T}(t)} = \sum_{t=1}^{T} \mathbf{1} \{ p_t \ge \lambda \} \gamma_{\mathcal{T}(t+1)-1} = \sum_{t=2}^{T+1} \mathbf{1} \{ p_{t-1} \ge \lambda \} \gamma_{\mathcal{T}(t)-1}.$$

Additionally, it is clear that $\mathcal{T}(\cdot)$ is a bijection mapping $\{1, 2 \leq t \leq T+1 : p_{t-1} \geq \lambda\}$ into $\{1, 2, \dots, \mathcal{T}(T+1)\}$. Hence, the latter sum can be rewritten as $\sum_{t=2}^{\mathcal{T}(T+1)} \gamma_{t-1} = \sum_{t=1}^{\mathcal{T}(T+1)-1} \gamma_t$ which provides (41).

Second, for proving (42), the crucial point is that according to the definition of $\mathcal{T}_j(T)$ (29), the functional $\mathcal{T}_j: \{\tau_j+1,\ldots\} \to \{1,\ldots\}$ is a bijection from $\{\tau_j+1\} \cup \{t \in \{\tau_j+2,\ldots,T+1\}: p_{t-1} \geq \lambda\}$ to $\{1,\ldots,\mathcal{T}_j(T+1)\}$, for any $j \geq 1$ and $T \geq \tau_j+1$. In particular, this entails

$$\begin{split} \sum_{p_t \geq \lambda, 1 \leq t \leq T} \gamma_{\mathcal{T}_j(t)} &= \sum_{p_t \geq \lambda, \tau_j + 1 \leq t \leq T} \gamma_{\mathcal{T}_j(t)} = \sum_{p_t \geq \lambda, \tau_j + 1 \leq t \leq T} \gamma_{\mathcal{T}_j(t+1) - 1} \\ &= \sum_{p_{t-1} \geq \lambda, \tau_j + 2 \leq t \leq T + 1} \gamma_{\mathcal{T}_j(t) - 1} = \sum_{t=2}^{\mathcal{T}_j(T+1)} \gamma_{t-1} = \sum_{t=1}^{\mathcal{T}_j(T+1) - 1} \gamma_t. \end{split}$$

This proves (42).

9. Online mFDR control for stopping times

Let us consider an almost surely finite stopping time, that is, any random variable τ taking values in $\{1, 2, \ldots\}$ with

- $\tau < +\infty$ almost surely;
- $\{\tau = t\} \in \mathcal{F}_t$ for all $t \geq 1$.

As a case in point, $\tau = \tau_r \wedge t_{\rm max}$ satisfies the two conditions above, where τ_r is the first time where the procedure makes at least a deterministic number $r \geq 1$ of rejections and $t_{\rm max} \geq 1$ is a deterministic maximum time for stopping. For some procedure \mathcal{A} , we define the τ -stopped mFDR as

$$\mathrm{mFDR}_{\tau}(\mathcal{A}, P) := \frac{\mathbf{E}_{X \sim P} (|\mathcal{H}_0 \cap \mathcal{R}(\tau)|)}{\mathbf{E}_{X \sim P} (1 \vee |\mathcal{R}(\tau)|)}, \tag{43}$$

similarly to the mFDR definition (6). The following result generalizes Lemma 4.1.

Lemma 9.1. Assume that both (2) and (3) hold, and consider a stopping time τ as above. For any procedure $\mathcal{A} = (\alpha_t, t \geq 1)$, if for some $\lambda \in [0, 1)$ we have $\sup_{T \geq 1} \widehat{\text{FDP}}_{\lambda}(T, \mathcal{A}) \leq \alpha$ a.s. (see (33)) then $\text{mFDR}_{\tau}(\mathcal{A}, P) \leq \alpha$.

Proof. We follow here Section 6.1 of Zrnic et al. (2021). Consider the process

$$M_t = \sum_{i \le t} \left(\mathbf{1} \{ p_i \le \alpha_i \} - \frac{\mathbf{1} \{ p_i > \lambda \}}{1 - \lambda} F_i(\alpha_i) \right), \quad t \ge 1.$$

By super-uniformity, $(M_t)_{t\geq 1}$ a super-martingale w.r.t. the filtration \mathcal{F}_t , that is, for all $t\geq 1$, M_t is measurable w.r.t. \mathcal{F}_t and $\mathbf{E}[M_{t+1}\,|\,\mathcal{F}_t]\leq M_t$. By the optional stopping theorem, we have $\mathbf{E}[M_{\tau}]\leq \mathbf{E}[M_1]\leq 0$. This implies

$$\mathbf{E}(|\mathcal{H}_0 \cap \mathcal{R}(\tau)|) = \mathbf{E}\left(\sum_{i \leq \tau, i \in \mathcal{H}_0} \mathbf{1}\{p_i \leq \alpha_i\}\right) \leq \mathbf{E}\left(\sum_{i=1}^{\tau} \frac{\mathbf{1}\{p_i > \lambda\}}{1 - \lambda} F_i(\alpha_i)\right)$$

$$\leq \mathbf{E}(\alpha(1 \vee R(\tau))),$$

where the last inequality follows from (33).

Theorem 9.1. In the setting of Theorem 4.3, for the procedure \mathcal{A} defined therein and for any stopping time as above, we also have $\operatorname{mFDR}_{\tau}(\mathcal{A}, P) \leq \alpha$ for all $P \in \mathcal{P}$.

Proof. The proof is straightforward from Lemma 9.1 and the proof of Theorem 4.3.

10. Online FDR control under super-uniformity

In this section, we study FDR control and not only mFDR control as in the main part of the paper (in a non-adaptive setting, or equivalently when $\lambda = 0$). We show that a sufficient condition for FDR control is independence and monotonicity, by slightly generalizing Theorem 2 of Ramdas et al. (2017) to the super-uniform setting of Section 2.1. We then show that $\mathcal{A}^{\rho \text{LORD}}$ does not satisfy the monotonicity condition in general, so that the FDR control of $\mathcal{A}^{\rho \text{LORD}}$ cannot be deduced. We then propose a monotone modification of $\mathcal{A}^{\rho \text{LORD}}$ that provably controls the FDR and which 'interpolates' between $\mathcal{A}^{\text{LORD}}$ and $\mathcal{A}^{\rho \text{LORD}}$.

10.1. General result

First define formally the FDR of a procedure \mathcal{A} for the parameter $P \in \mathcal{P}$ as

$$FDR(\mathcal{A}, P) := \sup_{T \ge 1} \{ FDR(T, \mathcal{A}, P) \}, \ FDR(T, \mathcal{A}, P) := \mathbf{E}_{X \sim P} \left(\frac{|\mathcal{H}_0 \cap \mathcal{R}(T)|}{1 \vee |\mathcal{R}(T)|} \right). \tag{44}$$

Theorem 10.1. Consider the setting of Section 2.1 where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. Consider any procedure $\mathcal{A} = \{\alpha_t, t \geq 1\}$ satisfying the two following conditions:

- (i) $\widehat{\text{FDP}}(T, \mathcal{A}) \leq \alpha$ for all $T \geq 1$ where $\widehat{\text{FDP}}(T, \mathcal{A})$ is the FDP estimate at time T defined by (26);
- (ii) the critical values of A are all nondecreasing with respect to the past rejections, that is, if the critical values α_T , $T \geq 1$, are written as a function $\alpha_T = \mathbf{A}_T(R_1, \ldots, R_{T-1})$ of the past rejection status $R_t = \mathbf{1}\{p_t \leq \alpha_t\}$, $t \leq T 1$, we have for all $T \geq 1$, $R_1, \ldots, R_{T-1}, R'_1, \ldots, R'_{T-1} \in \{0, 1\}$,

$$(\forall t \le T - 1, R'_t \ge R_t) \Rightarrow \mathbf{A}_T(R'_1, \dots, R'_{T-1}) \ge \mathbf{A}_T(R_1, \dots, R_{T-1}).$$
 (45)

Then, assuming that the model \mathcal{P} is such that (3) holds, we have $FDR(\mathcal{A}, P) \leq \alpha$.

Proof. We follow the proofs of Theorem 2 and Lemma 1 of Ramdas et al. (2017). Write $\alpha_T = \mathbf{A}_T(R_1, \dots, R_{T-1})$ as in statement (ii) of the theorem. By definition (44), and writing $R_t = \mathbf{1}\{p_t \leq \alpha_t\}, t \geq 1$, we have for $T \geq 1$,

$$FDR(T, \mathcal{A}, P) = \sum_{t \in \mathcal{H}_0, t \leq T} \mathbf{E} \left(\frac{\mathbf{1}\{p_t \leq \mathbf{A}_t(R_1, \dots, R_{t-1})\}\}}{1 \vee \sum_{t' \leq T} R_{t'}} \right)$$
$$= \sum_{t \in \mathcal{H}_0, t \leq T} \mathbf{E} \left(\frac{\mathbf{1}\{p_t \leq \mathbf{A}_t(R_1, \dots, R_{t-1})\}\}}{R_1 + \dots + R_{t-1} + 1 + R_{t+1} + \dots + R_T} \right),$$

because $R_t = 1$ on the event $\{p_t \leq \mathbf{A}_t(R_1, \dots, R_{t-1})\}$. Now, for a fixed $t \leq T$, let $\tilde{R}_{t'}^{(t)} = \mathbf{1}\{\tilde{p}_{t'}^{(t)} \leq \alpha_{t'}\}, t' \leq T$, where $\tilde{p}_{t'}^{(t)} = p_{t'}$ for $t' \neq t$ and $\tilde{p}_{t'}^{(t)} = 0$ otherwise. It is clear that we have $\mathbf{1}\{p_t \leq \mathbf{A}_t(R_1, \dots, R_{t-1})\} = \mathbf{1}\{p_t \leq \mathbf{A}_t(\tilde{R}_1^{(t)}, \dots, \tilde{R}_{t-1}^{(t)})\}$ and thus

$$\frac{\mathbf{1}\{p_t \le \mathbf{A}_t(R_1, \dots, R_{t-1})\}}{R_1 + \dots + R_{t-1} + 1 + R_{t+1} + \dots + R_T} = \frac{\mathbf{1}\{p_t \le \mathbf{A}_t(\tilde{R}_1^{(t)}, \dots, \tilde{R}_{t-1}^{(t)})\}}{\sum_{t' < T} \tilde{R}_{t'}^{(t)}},$$

because $\tilde{R}_{t'}^{(t)} = \tilde{R}_{t'}$ for all $t' \leq T$ provided that $R_t = 1$. Since by (3) p_t is independent of $p_{t'}$, $t' \neq t$, $t' \leq T$, and thus also of $(\tilde{R}_{t'}^{(t)}, t' \leq T)$, this gives

$$FDR(T, \mathcal{A}, P) = \sum_{t \in \mathcal{H}_0, t \leq T} \mathbf{E} \left(\frac{\mathbf{1}\{p_t \leq \mathbf{A}_t(\tilde{R}_1^{(t)}, \dots, \tilde{R}_{t-1}^{(t)})\}\}}{\sum_{t' \leq T} \tilde{R}_{t'}^{(t)}} \right)$$

$$= \sum_{t \in \mathcal{H}_0, t \leq T} \mathbf{E} \left(\frac{\mathbf{P}(p_t \leq \mathbf{A}_t(\tilde{R}_1^{(t)}, \dots, \tilde{R}_{t-1}^{(t)}) \mid p_{t'}, t' \leq T, t' \neq t)}{\sum_{t' \leq T} \tilde{R}_{t'}^{(t)}} \right)$$

$$\leq \sum_{t \in \mathcal{H}_0, t \leq T} \mathbf{E} \left(\frac{F_t(\mathbf{A}_t(\tilde{R}_1^{(t)}, \dots, \tilde{R}_{t-1}^{(t)}))}{\sum_{t' \leq T} \tilde{R}_{t'}^{(t)}} \right)$$

$$= \sum_{t \in \mathcal{H}_0, t \leq T} \mathbf{E} \left(\frac{F_t(\mathbf{A}_t(R_1, \dots, R_{t-1}))}{R_1 + \dots + R_{t-1} + 1 + \tilde{R}_{t+1}^{(t)} + \dots + \tilde{R}_T^{(t)}} \right),$$

by using in addition the super-uniformity (2) in the third line. Now, $\tilde{R}_{t}^{(t)} = 1 \geq R_{t}$ and by (45), we have $\tilde{R}_{t+1}^{(t)} = \mathbf{1}\{\tilde{p}_{t+1}^{(t)} \leq \mathbf{A}_{t+1}(R_{1}, \dots, R_{t-1}, 1)\} = \mathbf{1}\{p_{t+1} \leq \mathbf{A}_{t+1}(R_{1}, \dots, R_{t-1}, R_{t})\} = R_{t+1},$ so by induction we have $\tilde{R}_{t'}^{(t)} \geq R_{t'}$ for all $t' \geq t$. This yields

$$FDR(T, \mathcal{A}, P) \le \sum_{t \in \mathcal{H}_0, t \le T} \mathbf{E}\left(\frac{F_t(\alpha_t)}{R(T) \vee 1}\right) \le \alpha,$$

because by assumption $\widehat{\text{FDP}}(T, \mathcal{A}) \leq \alpha$, see (26).

Unfortunately, in general, $\mathcal{A}^{\rho \text{LORD}}$ does not satisfy (45).

Proposition 10.1. There exists a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ such that the 'greedy' critical values (27) of $\mathcal{A}^{\rho LORD}$ do not satisfy (45) pointwise.

Proof. Let us consider a sequence $(R_1, \ldots, R_5) = (0, 0, 1, 0, 1)$ so that $\tau_1 = 3$ and $\tau_2 = 5$, then the corresponding critical values (27) are given by

- $\bullet \ \alpha_1 = W_0 \gamma_1$
- $\alpha_2 = W_0 \gamma_2 + \alpha_1 F_1(\alpha_1)$
- $\bullet \ \alpha_3 = W_0 \gamma_3 + \alpha_2 F_2(\alpha_2)$
- $\alpha_4 = W_0 \gamma_4 + (\alpha W_0) \gamma_1 + \alpha_3 F_3(\alpha_3)$
- $\alpha_5 = W_0 \gamma_5 + (\alpha W_0) \gamma_2 + \alpha_4 F_4(\alpha_4)$
- $\alpha_6 = W_0 \gamma_6 + (\alpha W_0) \gamma_3 + \alpha \gamma_1 + \alpha_5 F_5(\alpha_5)$

Let us consider a sequence $(R'_1,\ldots,R'_5)=(0,1,1,0,1)$ so that $\tau'_1=2,\,\tau'_2=3$ and $\tau_3' = 5$, then the corresponding critical values (27) are given by

- $\bullet \ \alpha_1' = W_0 \gamma_1$

- $\alpha_1 W_0 \gamma_1$ $\alpha_2' = W_0 \gamma_2 + \alpha_1' F_1(\alpha_1')$ $\alpha_3' = W_0 \gamma_3 + (\alpha W_0) \gamma_1 + \alpha_2' F_2(\alpha_2')$ $\alpha_4' = W_0 \gamma_4 + (\alpha W_0) \gamma_2 + \alpha \gamma_1 + \alpha_3' F_3(\alpha_3')$ $\alpha_5' = W_0 \gamma_5 + (\alpha W_0) \gamma_3 + \alpha \gamma_2 + \alpha_4' F_4(\alpha_4')$ $\alpha_6' = W_0 \gamma_6 + (\alpha W_0) \gamma_4 + \alpha \gamma_3 + \alpha \gamma_1 + \alpha_5' F_5(\alpha_5')$.

Hence, $\alpha_1 = \alpha_1'$, $\alpha_2 = \alpha_2'$, $\alpha_3' > \alpha_3$, and

$$\alpha'_{4} = W_{0}\gamma_{4} + (\alpha - W_{0})\gamma_{2} + \alpha\gamma_{1} + \alpha'_{3} - F_{3}(\alpha'_{3})$$

$$= \alpha_{4} + (\alpha - W_{0})\gamma_{2} + \alpha\gamma_{1} - (\alpha - W_{0})\gamma_{1} + \alpha'_{3} - \alpha_{3} + F_{3}(\alpha_{3}) - F_{3}(\alpha'_{3})$$

$$= \alpha_{4} + (\alpha - W_{0})\gamma_{2} + \alpha\gamma_{1} + F_{3}(\alpha_{3}) - F_{3}(\alpha'_{3}).$$

In particular, if $F_3(\alpha_3) = \alpha_3'$ (minimal super-uniformity reward) while $F_3(\alpha_3) =$ 0 (maximal super-uniformity reward) and $F_1(\alpha'_1) = F_2(\alpha'_2) = 0$, we have

$$\alpha_3' = W_0(\gamma_1 + \gamma_2 + \gamma_3) + (\alpha - W_0)\gamma_1$$

and thus

$$\alpha_4' = \alpha_4 + (\alpha - W_0)\gamma_2 + \alpha\gamma_1 - W_0(\gamma_1 + \gamma_2 + \gamma_3) - (\alpha - W_0)\gamma_1.$$

As a result, for $W_0 = \alpha/2$, $\alpha'_4 = \alpha_4 - W_0 \gamma_3 < \alpha_4$. Finally, with the notation of condition (45), we have for T=4, that $(\forall t \leq T-1, R'_t \geq R_t)$ but $\alpha'_T=$ $\mathbf{A}_T(R_1',\ldots,R_{T-1}') \geq \mathbf{A}_T(R_1,\ldots,R_{T-1}) = \alpha_T$, which shows that condition (45) is violated.

10.2. A SUR procedure controlling the FDR

We propose the following modification of $\mathcal{A}^{\rho LORD}$.

Definition 10.1. For a spending sequence γ and a SUR spending sequence γ' , the modified LORD procedure with super-uniformity reward, denoted by $\mathcal{A}^{\widetilde{\rho LORD}} = \{\alpha_t^{\widetilde{\rho LORD}}, t \geq 1\}$, is defined by the following recursion on $T \geq 1$: given critical values $\alpha_1^{\widetilde{\rho LORD}}, \ldots, \alpha_{T-1}^{\widetilde{\rho LORD}}$, put $\alpha_T^{\widetilde{\rho LORD}} = \mathbf{A}_T^{\widetilde{\rho LORD}}(R_1, \ldots, R_{T-1})$ with

$$\mathbf{A}_{T}^{\widetilde{\rho_{LORD}}}(R_{1},\ldots,R_{T-1}) = \mathbf{A}_{T}^{LORD}(R_{1}^{0},\ldots,R_{T-1}^{0}) + \min_{\substack{(r_{1},\ldots,r_{T-1})\in\{0,1\}^{T-1}\\\forall i\leq T-1,\ r_{i}\geq R_{i}}} \left(\sum_{t=1}^{T-1} \gamma_{T-t}'\rho_{t}(r_{1},\ldots,r_{t-1})\right),$$
(46)

where $\mathbf{A}_{T}^{LORD}(R_{1}^{0},\ldots,R_{T-1}^{0}) = \alpha_{T}^{LORD}$ is given by (25) with $R_{t}^{0} = \mathbf{1}\{p_{t} \leq \alpha_{t}^{LORD}\}$, $R_{t} = \mathbf{1}\{p_{t} \leq \alpha_{t}^{\widehat{\rho LORD}}\}$ and the super-uniformity reward at time t for $\widehat{\mathcal{A}^{\rho LORD}}$ is given by $\rho_{t}(r_{1},\ldots,r_{t-1}) = \widehat{\mathbf{A}_{t}^{\widehat{\rho LORD}}}(r_{1},\ldots,r_{t-1}) - F_{t}(\widehat{\mathbf{A}_{t}^{\widehat{\rho LORD}}}(r_{1},\ldots,r_{t-1}))$ (for the past rejections r_{1},\ldots,r_{t-1}).

First note that when $F_t(x) = x$, then $\mathcal{A}^{\rho LORD}$ reduces to the LORD procedure because the min-component in (46) disappears. Otherwise, the procedure provides some reward. Compared to $\mathcal{A}^{\rho LORD}$, the modification $\mathcal{A}^{\rho LORD}$ involves a "least favorable" super-uniformity reward over all possible past rejection sequences that could have occurred in the past. Computing these critical values becomes intractable when t gets large because of the minimum which requires to compute all $\alpha_t^{\rho LORD}(r_1,\ldots,r_{t-1})$, for all possible rejection sequences $(r_1,\ldots,r_{T-1}) \in \{0,1\}^{t-1}$. Also, $\alpha_t^{\rho LORD}$ is predictable in the sense of the present theory — and hence defines a valid online procedure — only after enlarging the filtration \mathcal{F}_{t-1} to contain R_1^0,\ldots,R_{T-1}^0 the sequence of rejections for the standard LORD procedure, that should be run in parallel when computing (46). The counterpart of these drawbacks is that $\mathcal{A}^{\rho LORD}$ controls the online (non marginal) FDR, as the next result establishes.

Theorem 10.2. Consider the setting of Section 2.1 where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. Consider $\mathcal{A}^{\widetilde{\rho LORD}} = \{\alpha_t^{\widetilde{\rho LORD}}, t \geq 1\}$ given in Definition 10.1. Then we have

$$\forall T \ge 1, \ \alpha_T^{LORD} \le \widetilde{\alpha_T^{\rho LORD}} \le \alpha_T^{\rho LORD},$$
 (47)

and $\widetilde{\mathcal{A}^{\rho LORD}}$ satisfies conditions (i) and (ii) of Theorem 10.1. In particular, assuming that the model \mathcal{P} is such that (3) holds, we have $\mathrm{FDR}(\widetilde{\mathcal{A}^{\rho LORD}}, P) \leq \alpha$. Proof. First, since

$$0 \leq \min_{\substack{(r_1, \dots, r_{T-1}) \in \{0,1\}^{t-1} \\ \forall i \leq T-1, r_i \geq R_i}} \left(\sum_{t=1}^{T-1} \gamma'_{T-t} \rho_t(r_1, \dots, r_{t-1}) \right) \leq \sum_{t=1}^{T-1} \gamma'_{T-t} \rho_t(R_1, \dots, R_{t-1})$$

relation (47) easily follows from (46). Second, this minimum is clearly nondecreasing w.r.t. (R_1, \ldots, R_{T-1}) , so that by definition (46) $\mathcal{A}^{\rho LORD}$ satisfies (45).

Now let us check condition (i) of Theorem 10.1. For this we consider the new critical values sequence (slightly different than $\alpha^{\rho \text{LORD}}$) given by

$$\alpha_T = \mathbf{A}_T^{\text{LORD}}(R_1^0, \dots, R_{T-1}^0) + \sum_{t=1}^{T-1} \gamma'_{T-t} \rho_t(R_1, \dots, R_{t-1}),$$

where $R_t^0 = \mathbf{1}\{p_t \leq \alpha_t^{\text{LORD}}\}, R_t = \mathbf{1}\{p_t \leq \alpha_t\}$. The same reasoning as above gives the domination $\forall T \geq 1, \alpha_T^{\widehat{\text{PLORD}}} \leq \alpha_T$. As a result, we have

$$\alpha_T^{\widetilde{\rho_{\text{LORD}}}} + \sum_{t=1}^{T-1} F_t(\alpha_t^{\widetilde{\rho_{\text{LORD}}}}) \leq \alpha_T^{\widetilde{\rho_{\text{LORD}}}} + \sum_{t=1}^{T-1} \left[(1 - a_{T-t}) \alpha_t^{\widetilde{\rho_{\text{LORD}}}} + a_{T-t} F_t(\alpha_t^{\widetilde{\rho_{\text{LORD}}}}) \right]$$

$$\leq \alpha_T + \sum_{t=1}^{T-1} \left[(1 - a_{T-t}) \alpha_t + a_{T-t} F_t(\alpha_t) \right].$$

Now applying Lemma 8.2 with $\alpha_T^0 = \mathbf{A}_T^{\text{LORD}}(R_1^0,\dots,R_{T-1}^0)$ and $\lambda=0$, the last display is equal to $\alpha_T^0 + \sum_{t=1}^{T-1} \mathbf{1}\{p_t \geq \lambda\}\alpha_t^0 \leq \alpha(1 \vee R^0(T))$. But now the latter is smaller than $\alpha\left(1 \vee \sum_{t \leq T} \mathbf{1}\{p_t \leq \alpha_t^{\rho \widehat{\text{LORD}}}\}\right)$ by using (47) (left inequality). This means that $\widehat{\text{FDP}}(T, \mathcal{A}^{\rho \widehat{\text{LORD}}}) \leq \alpha$ holds and shows that condition (i) of Theorem 10.1 is satisfied. The FDR control hence follows from Theorem 10.1.

11. Delayed spending approach

In this section we present another way of incorporating super-uniformity into OMT which we refer to as *delayed spending* (in the sequel abbreviated as DS). We are grateful to Aaditya Ramdas for this suggestion.

11.1. Motivation

In the discrete setting, let us consider the following constrained spending problem: at each step t, choose the critical value α_t to be in the support S_t (including 0) such that the following constraint holds

$$\sum_{t>1} \alpha_t \le \alpha. \tag{48}$$

It solves the super-uniformity problem, because $F_t(\alpha_t) = \alpha_t$ for all t, while it controls the online FWER. This general principle, that we refer to as 'constrained spending strategies', can be implemented in many ways.

Markedly, the SUR approach is a way to achieve this, by additionally following some reference critical values — here the online Bonferroni critical values α_t^{OB} (10). Indeed, the rejection decision $p_t \leq \alpha_t^{\rho^{\text{OB}}}$ and $p_t \leq \alpha_t = F_t(\alpha_t^{\rho^{\text{OB}}})$ are almost surely identical and we have calibrated $\alpha_t^{\rho^{\text{OB}}}$ such that (48) holds,

see (13). In other words, even if our critical values are not constrained to be in the support initially, the effective critical values $\alpha_t = F_t(\alpha_t^{\rho \text{OB}})$ that are actually used in the decision rule will automatically belong to the support. Thus, our approach can be equivalently seen as a way of implementing the constrained spending strategies delineated above.

Obviously, there are other ways to implement the constrained spending strategy. One instance is the delayed spending (DS) approach, that we describe below.

The new procedure is introduced in Section 11.2, while we highlight some mathematical and practical differences with our approach in Sections 11.3 and 11.4. In order to make the new procedure more efficient we also present a hybrid version in Section 11.5. For simplicity, we restrict ourselves to FWER controlling procedures for discrete data throughout this section.

11.2. Definition

Let us start with the critical value $\alpha_1 = \alpha \gamma_1$. While the OB procedure would choose $\alpha_2 = \alpha \gamma_2$, the idea is that if the super-uniformity is strong enough to ensure $F_1(\alpha \gamma_1) + F_2(\alpha \gamma_1) \leq \alpha \gamma_1$, we can still use $\alpha_2 = \alpha \gamma_1$ in the second round. This process can be continued until $F_1(\alpha \gamma_1) + \cdots + F_{b_1+1}(\alpha \gamma_1) > \alpha \gamma_1$, in which case we switch to $\alpha_{b_1} = \alpha \gamma_2$, and so on. This way, we can incorporate the super-uniformity directly by 'delaying' the γ sequence.

More formally, consider the setting of Section 2.1, where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (2) is at hand. The above strategy reads:

$$\alpha_t^{\text{DS}} = \alpha \gamma_{\mathcal{C}(t)}, \text{ where } \mathcal{C}(t) = \min\{j \ge 1 : b_j \ge t\}, \ t \ge 1;$$
 (49)

$$b_j = \max \left\{ T \ge b_{j-1} + 1 : \sum_{t=b_{j-1}+1}^T F_t(\alpha \gamma_j) \le \alpha \gamma_j \right\}, \quad j \ge 1,$$
 (50)

(with the convention $b_0 = 0$ and $b_j = +\infty$ if the set in (50) is empty), so that $j = \mathcal{C}(t)$ for $b_{j-1} + 1 \le t \le b_j$. Thus, the DS method processes each sub-budget $\alpha \gamma_j$ one at a time, until the stopping rule in (50) is met and the transition to the next sub-budget $\alpha \gamma_{j+1}$ is made. Since $\mathcal{C}(t) \le t$ we can interpret $\alpha_t^{\mathrm{DS}} = \alpha \gamma_{\mathcal{C}(t)}$ as a 'slowed-down' variant of the original OB procedure.

The procedure (49) controls the online FWER under (2) because by (13), a sufficient condition is given by $\sum_{t=1}^{T} F_t(\alpha_t) \leq \alpha$, $T \geq 1$, and we indeed have

$$\sum_{t\geq 1} F_t(\alpha_t^{\text{DS}}) \leq \sum_{j\geq 1} \sum_{t=b_{j-1}+1}^{b_j} F_t(\alpha \gamma_j) \leq \sum_{j\geq 1} \alpha \gamma_j \leq \alpha,$$

by definition of the b_j 's. Note that the b_j 's are based on local averages (in time) of the $F_t(x)$'s at certain points x. This shares similarity to the approach of Westfall and Wolfinger (1997) for offline FWER control.

11.3. Comparison to SUR for real data

Both the DS and the SUR approaches use super-uniform rewarding. In a nutshell, the DS approach slows down the clock whereas the SUR approach augments the critical values of existing OMT procedures in an additive way. While a more detailed comparison can be found in the following Section 11.4, we may say that no method dominates the other one uniformly. The examples given in Section 11.4 (delayed start, long/infinite delay, ineffective delay) suggest that the DS method could be more efficient at the very start of the stream but may suffer from conservativeness afterwards.

To assess the behaviour of the procedures in a practical setting, we reanalyse the IMPC data from Section 5.3 using the DS procedure defined by (49) and (50) and compare it with the OB and ρ OB from Section 3. The results for FWER control at level $\alpha = 0.2$ are displayed in Table 5 and Figure 9. As Figure 9 (right panel) shows, the rejection process $\{R(T), T \geq 1\}$, is almost identical at the very start. However, for larger T, the delayed approach makes less discoveries than the ρ OB procedure and this, uniformly in time for this data set. This conservative behaviour is probably caused by under-utilization of wealth as described in Section 11.4. More specifically, the non-utilized component of $\alpha = 0.2$ accumulates up to time T = 1500 approximately to 0.077, so that approximately 38.5% of $\alpha = 0.2$ are effectively neglected. Accordingly, the wealth plot displayed in Figure 9 shows that the delayed approach manages

Table 5 Number of discoveries for SURE online Bonferroni (15) (bandwidth h = 10) and the DS

approach (49). Here $C(30\,000) = 5083$ as defined in (49). These numbers are obtained by running the procedures on the first 30 000 genes for male (second row) and female (third row) mice in the IMPC data.

Procedures	OB	ρOB	Delayed
# discoveries (male)	229	377	293
# discoveries (female)	267	481	355

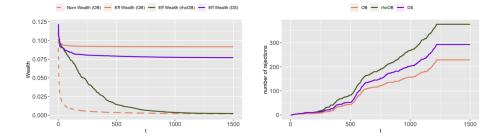


Fig 9. Comparison with DS. Left: nominal wealth for OB (dashed orange curve), effective wealth for OB (solid orange curve), effective wealth for ρ OB (solid green curve) and effective wealth for DS (solid purple curve), plot similar to Figure 2. Right: rejection numbers, cumulated over time, for the same procedures (same color code). Both plots are computed from the male IMPC data.

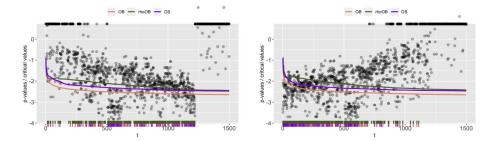


Fig 10. Critical values of OB (orange), ρ OB (green) and DS (purple) for the IMPC data (left panel is for male, right panel is for female).

to spend more wealth than the OB procedure, but still deviates strongly from the nominal wealth curve. Figure 10 illustrates the same phenomenon for the critical values.

11.4. Formal properties

From the definition of the DS approach we obtain the following comparison to OB and ρ OB:

- the DS approach improves OB uniformly when γ_t is nonincreasing: indeed $\mathcal{C}(t) \leq t$, so that $\alpha_t^{\text{DS}} = \alpha \gamma_{\mathcal{C}(t)} \geq \alpha \gamma_t = \alpha_t^{\text{OB}}$.
- the DS approach does not depend on any other tuning parameter such as the bandwidth. By contrast, choosing this parameter badly in the ρ OB procedure may adversely affect its performance.
- the DS approach is another way of using the super uniformity reward. For instance, if there is no super uniformity reward, that is, $F_t(\alpha \gamma_t) = \alpha \gamma_t$ for all t, then $b_t = t$ and the DS procedure reduces to OB.

In addition, we have the following observations:

- Delayed start: If $F_t(x) = 0$ for all x < 1 and $t \le T_0$ and $F_t(x) = x$ for $t \ge T_0 + 1$, the DS procedure is much more intuitive: it yields $b_1 = T_0 + 1$ by (50) and $\alpha_t^{\text{DS}} = \alpha \gamma_{t-T_0}$ for $t \ge T_0 + 1$ which is the most natural way to proceed (just start the testing process at time $T_0 + 1$). By contrast, ρ OB (with rectangular kernel of bandwidth r) collects some reward in $\alpha_t^{\rho \text{OB}}$, $1 \le t \le T_0$, spends the reward in the following r time points, but continues with $\alpha_t^{\rho \text{OB}} = \alpha \gamma_t$ for $t \ge T_0 + r + 1$. Hence, delaying spends the super-uniformity more intuitively than ρ OB in that situation. More generally, in practice, we may therefore expect DS to be more efficient in the beginning of the stream.
- Long/infinite delay: Conversely, if there exists $T_0 \geq 1$ such that for all $t \geq b_{T_0}+1$, $F_t(\alpha\gamma_{\mathcal{C}(T_0)+1})=0$, then we have $b_{T_0+1}=+\infty$ from (50), which in turn implies $\mathcal{C}(t) \leq T_0+1$. But for $t \geq b_{T_0}+1$, we have $\mathcal{C}(t) \geq T_0+1$ by (49). Hence, for $t \geq b_{T_0}+1$, $\mathcal{C}(t)=T_0+1$ and the 'spending clock'

freezes. On the one hand, we have $\alpha_t^{\mathrm{DS}} = \alpha \gamma_{T_0+1}$ so the delaying works perfectly to effectively improve the OB critical values. On the other hand, this effectively stops the spending of any further budget and thus a large part of the wealth is left unspent. This is in contrast to the SUR approach which uses a reward of an additive nature and thus always has a chance to spend the budget.

• Under-utilization of wealth. The DS method processes each sub-budget $\alpha\gamma_j$ one at a time, until the transition to the next sub-budget $\alpha\gamma_{j+1}$ is made. In most cases, however, the inequality (50) defining the transition time b_j will be a strict inequality, meaning that when we move on to the next sub-budget we will have used $\sum_{t=b_{j-1}+1}^{b_j} F_t(\alpha\gamma_j) < \alpha\gamma_j$. Thus, this method does not exhaust the available sub-budgets. Moreover, since it neglects these 'alpha-gaps', they accumulate over time. This under-utilized wealth leads to unnecessary conservatism. Removing such gaps was precisely the primary motivation for introducing our SUR method, see Section 2.3.

The most disadvantageous scenario occurs when $b_t = t$ for all $t \leq T$, so that the DS procedure reduces to the original OB procedure up to time T. As an example consider $\epsilon \in (0, \alpha \gamma_T)$ for some large $T \geq 1$ and assume that the support of each p_t is given by $S_t = \{\epsilon, A_t, \alpha \gamma_{t-1}\} \cup \{1\}$ (convention $\alpha \gamma_0 = 1$), where A_t is a finite subset of $(\alpha \gamma_t, \alpha \gamma_{t-1})$. Then we have $F_1(\alpha \gamma_1) + F_2(\alpha \gamma_1) = \alpha \gamma_1 + \epsilon$ hence $b_1 = 1$, and more generally $F_t(\alpha \gamma_t) + F_{t+1}(\alpha \gamma_t) = \alpha \gamma_t + \epsilon$ for all $t \leq T$, which implies $b_t = t$ for all $t \leq T$. However, we know that OB does not allow to spend all the budget in such a discrete situation, see Figure 2.

A potential remedy for the conservatism of the DS method could be to combine it with our SUR method. We describe such a hybrid approach in more detail in Section 11.5.

In summary, it may be said that the delaying method is particularly appealing in terms of simplicity and elegance, while the primary aim of the SUR approach is on efficiency.

11.5. Hybrid approach

In this section, we describe a hybrid approach, combining the ideas underlying DS and SUR, in order to improve the utilization of wealth of DS.

The method starts as follows: first let $\alpha_1^{\text{Hyb}} = \alpha \gamma_1, \ldots, \alpha_{b_1}^{\text{Hyb}} = \alpha \gamma_1$ as long as $F_1(\alpha \gamma_1) + \cdots + F_{b_1}(\alpha \gamma_1) \leq \alpha \gamma_1$. Then consider the reward $\rho_1 = \alpha \gamma_1 - (F_1(\alpha \gamma_1) + \cdots + F_{b_1}(\alpha \gamma_1))$ and let $\alpha_{b_1+1}^{\text{Hyb}} = \alpha \gamma_2 + \rho_1, \ldots, \alpha_{b_2}^{\text{Hyb}} = \alpha \gamma_2 + \rho_1$ as long as $F_{b_1+1}(\alpha \gamma_2 + \rho_1) + \cdots + F_{b_2}(\alpha \gamma_2 + \rho_1) \leq \alpha \gamma_2 + \rho_1$. More generally, let $b_0 = 0, \ \rho_0 = 0$, and for all $j \geq 1$,

$$\alpha_{b_{j-1}+1}^{\text{\tiny Hyb}} = \alpha \gamma_j + \rho_{j-1}, \dots, \alpha_{b_j}^{\text{\tiny Hyb}} = \alpha \gamma_j + \rho_{j-1}$$

$$b_{j} = \max \left\{ T \ge 1 : \sum_{t=b_{j-1}+1}^{T} F_{t}(\alpha \gamma_{j} + \rho_{j-1}) \le \alpha \gamma_{j} + \rho_{j-1} \right\}$$
$$\rho_{j} = \alpha \gamma_{j} + \rho_{j-1} - \left(\sum_{t=b_{j-1}+1}^{b_{j}} F_{t}(\alpha \gamma_{j} + \rho_{j-1}) \right).$$

Then the online FWER control holds because for all $j_0 \geq 1$, we have

$$\sum_{t\geq 1} F_t(\alpha_t^{\text{Hyb}}) = \sum_{j=1}^{j_0} \left(\sum_{t=b_{j-1}+1}^{b_j} F_t(\alpha \gamma_j + \rho_{j-1}) \right) + F_{b_{j_0}+1}(\alpha \gamma_{j_0+1} + \rho_{j_0})$$

$$\leq \sum_{j=1}^{j_0} (\alpha \gamma_j + \rho_{j-1} - \rho_j) + \alpha \gamma_{j_0+1} + \rho_{j_0} = \sum_{j=1}^{j_0+1} \alpha \gamma_j \leq \alpha,$$

because $\sum_{j=1}^{j_0} (\rho_{j-1} - \rho_j) = -\rho_{j_0}$ (telescopic sum). When $\rho_t = 0$ for all $t \ge 1$, the hybrid approach reduces to the DS approach. When $b_j = j$, the hybrid approach reduces to the greedy SUR procedure.

We can also combine the DS with smoothed SUR rewarding, which gives us the following, slightly more involved, procedure. For some SUR spending sequence $\gamma' = (\gamma_t')_{t\geq 1}$ (nonnegative and such that $\sum_{t\geq 1} \gamma_t' \leq 1$), let $b_0 = 0$, $\rho_0 = 0$ and for all $j \geq 1$,

$$\alpha_{b_{j-1}+1}^{\text{Hyb}} = \alpha \gamma_{j} + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_{i}, \dots, \quad \alpha_{b_{j}}^{\text{Hyb}} = \alpha \gamma_{j} + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_{i}$$

$$b_{j} = \max \left\{ T \ge 1 : \sum_{t=b_{j-1}+1}^{T} F_{t} \left(\alpha \gamma_{j} + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_{i} \right) \le \alpha \gamma_{j} + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_{i} \right\}$$

$$\rho_{j} = \alpha \gamma_{j} + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_{i} - \left(\sum_{t=b_{j-1}+1}^{b_{j}} F_{t} \left(\alpha \gamma_{j} + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_{i} \right) \right).$$

The online FWER control holds because for all $j_0 \geq 1$, we have

$$\sum_{t \ge 1} F_t(\alpha_t^{\text{Hyb}}) \le \sum_{j=1}^{j_0} \left(\sum_{t=b_{j-1}+1}^{b_j} F_t \left(\alpha \gamma_j + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_i \right) \right) + \alpha \gamma_{j_0+1} + \sum_{i=1}^{j_0} \gamma'_{j_0+1-i} \rho_i.$$

Now letting $a_T = \sum_{t=1}^T \gamma_t'$, we obtain

$$\sum_{j=1}^{j_0} \left(\sum_{t=b_{j-1}+1}^{b_j} F_t \left(\alpha \gamma_j + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_i \right) \right) \\
\leq \sum_{j=1}^{j_0} a_{j_0-j+1} \left(\alpha \gamma_j + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_i - \rho_j \right) + \sum_{j=1}^{j_0} (1 - a_{j_0-j+1}) \left(\alpha \gamma_j + \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_i \right) \\$$

$$= \sum_{j=1}^{j_0} \alpha \gamma_j + \sum_{j=1}^{j_0} \sum_{i=1}^{j-1} \gamma'_{j-i} \rho_i - \sum_{j=1}^{j_0} a_{j_0-j+1} \rho_j$$
$$= \sum_{j=1}^{j_0} \alpha \gamma_j + \sum_{i=1}^{j_0-1} a_{j_0-i} \rho_i - \sum_{j=1}^{j_0} a_{j_0-j+1} \rho_j,$$

and the latter is equal to $\sum_{j=1}^{j_0} \alpha \gamma_j - \sum_{j=1}^{j_0-1} \gamma'_{j_0-j+1} \rho_j - a_1 \rho_{j_0} = \sum_{j=1}^{j_0} \alpha \gamma_j - \sum_{j=1}^{j_0} \gamma'_{j_0-j+1} \rho_j$. Combining this with the above bound for the FWER concludes the proof.

To compare the performance of the hybrid approach with the SUR and DS approaches, we use the simulation setting from Section 5.2 in the case where the signal is positioned at the beginning of the stream for each simulation run, which is the most favorable position of the signal for any procedure (see Section 5.2.2 for more details). We consider both procedures based on the uniform kernel (bandwidth h=100) and those based on the greedy spending sequence (denoted by 'greedy').

Figure 11 shows that taking super-uniformity into account is always beneficial, regardless of the specific approach used. The base DS method performs similarly to the greedy ρ OB and the greedy hybrid. In contrast, the hybrid approach based on a uniform kernel improves DS, with performance close to ρ OB. Hence, we conclude that closing the alpha-gaps by smoothing with an adequate kernel can make the hybrid approach as powerful as the smoothed ρ OB method. However, given the added complexity of the hybrid approach, we prefer to stick with the smoothed SUR.

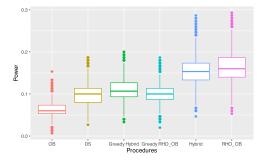


FIG 11. Power of several online FWER controlling approaches for simulated data (see text): online Bonferroni (OB), Delayed spending (DS), greedy hybrid, greedy ρ OB, hybrid, ρ OB.

12. Complements on generalized α -investing rules

12.1. SUR-GAI++ rules

GAI++ rules have been introduced in Ramdas et al. (2017) to control the (m)FDR. Here, we can extend them to our super-uniform setting as follows. Let

us consider the following recursive constraints: for $t \geq 1$,

$$R_t = \mathbf{1}\{p_t \le \alpha_t\}$$

$$W(t) = W(t-1) - \phi_t + R_t \psi_t \text{ 'wealth available at time } t+1\text{'}$$

$$\phi_t \in [0, W(t-1)] \text{ 'spent at time } t\text{'}$$

$$\psi_t \le b_t + \min(\phi_t, \phi_t / F_t(\alpha_t) - 1) \text{ 'reward at time } t\text{'}$$

$$\psi_t \ge 0$$

$$b_t = \alpha - W_0 \mathbf{1}\{t \le \tau_1\},$$

where $W(0) = W_0 \in [0, \alpha]$. Any choice of W_0 and α_t, ϕ_t, ψ_t that are \mathcal{F}_{t-1} measurable and satisfying the above constraints defines a SUR-GAI++ procedure. Here, the only difference with the original GAI++ rule is the presence of $F_t(\alpha_t)$ instead of α_t in the definition of Ψ_t .

Proposition 12.1. Consider the setting of Section 2.1 where a null bounding family $\mathfrak{F} = \{F_t, t \geq 1\}$ satisfying (4) is at hand. Then any SUR-GAI++ procedure controls the mFDR at level α .

The proof is totally analogous to the one of Theorem 1 in Ramdas et al. (2017) (adapted to the mFDR, so without using any monotonicity property).

12.2. GAI++ weighting

Consider (continuous) p-values satisfying (1)-(3) and weights $w_t \geq 0$ that are \mathcal{F}_{t-1} measurable for all t. In Section 5 of Ramdas et al. (2017), the following (implicit) GAI++ weighting scheme has been proposed:

$$R_t = \mathbf{1}\{p_t \le w_t \alpha_t\}$$

$$W(t) = W(t-1) - \phi_t + R_t \psi_t$$

$$\phi_t \in [0, W(t-1)]$$

$$\psi_t \le b_t + \min(\phi_t, \phi_t / (w_t \alpha_t) - 1)$$

$$\psi_t \ge 0$$

$$b_t = \alpha - W_0 \mathbf{1}\{t < \tau_1\}.$$

Note that the latter constraints are similar to the constraints given in Section 12.1 for $F_t(x) = (w_t x) \wedge 1$ (up to the '\lambda1' which makes the constraints here slightly more stringent) so that this weighting case is a particular SUR-GAI++ procedure.

For given raw weights $r_t \ge 0$ (\mathcal{F}_{t-1} measurable), an explicit procedure which is used in Ramdas et al. $(2017)^6$, is obtained by choosing α_t , w_t , ϕ_t , ψ_t as follows:

$$w_t = r_t \wedge \frac{1}{1 - b_t}$$

 $^{^6\}mathrm{This}$ procedure is available at https://github.com/fanny-yang/OnlineFDRCode

$$\phi_t = \alpha_t = W_0 \gamma_t + \sum_{j \ge 1} \gamma_{t-\tau_j} \psi_{\tau_j}$$

$$\psi_t = b_t + \min \left(\phi_t, 1/w_t - 1 \right).$$

This choice is valid because $\alpha_t \leq W(t-1)$ for all t. Indeed,

$$W(t-1) = W_0 + \sum_{i=1}^{t-1} (-\alpha_i + R_i \psi_i),$$

so $\alpha_t \leq W(t-1)$ if and only if $\sum_{i=1}^t \alpha_i \leq W_0 + \sum_{i=1}^{t-1} R_i \psi_i$, which is true.

12.3. Our ρ-LORD is a SUR-GAI++ rule

We claim here that the procedure ρ -LORD corresponds to a SUR-GAI++ rule with the choice $\phi_t = F_t(\alpha_t)$, $\psi_t = b_t$, and

$$\alpha_t = W_0 \gamma_t + (\alpha - W_0) \gamma_{t-\tau_1} + \alpha \sum_{j>2} \gamma_{t-\tau_j} + \sum_{i=1}^{t-1} \gamma'_{t-i} \rho_i \quad t \ge 1.$$
 (51)

To establish this, we check that all constraints given in Section 12.1 are satisfied. The only non-trivial one is $\phi_t = F_t(\alpha_t) \leq W(t-1)$. Let us now prove it. Recall that $W(t) = W(t-1) - \phi_t + R_t b_t$ and $W(0) = W_0$. Hence $\alpha_1 = W_0 \gamma_1 \leq W_0$. Moreover, for $t \geq 2$,

$$W(t-1) = W_0 + (\alpha - W_0)\mathbf{1}\{t-1 \ge \tau_1\} + \alpha \sum_{j \ge 2} \mathbf{1}\{t-1 \ge \tau_j\} - \sum_{i=1}^{t-1} F_i(\alpha_i).$$

So we have $\bar{\alpha}_t \leq W(t-1)$ for the critical value

$$\bar{\alpha}_{t} = \left(\sum_{i=1}^{t} \gamma_{i}\right) W_{0} + \sum_{i=1}^{t-1} \left((\alpha - W_{0}) \gamma_{i-\tau_{1}+1} \mathbf{1} \{ i \geq \tau_{1} \} + \alpha \sum_{j \geq 2} \gamma_{i-\tau_{j}+1} \mathbf{1} \{ i \geq \tau_{j} \} \right)$$
$$- \sum_{i=1}^{t-1} \left[a_{t-i} F_{i}(\bar{\alpha}_{i}) + (1 - a_{t-i}) \bar{\alpha}_{i} \right],$$

by letting $a_t = \sum_{i=1}^t \gamma_i'$. But now, we have that $\bar{\alpha}_t = \alpha_t$ for all t, for α_t defined by (51). Indeed, this can be seen from Lemma 8.2, applied with $\lambda = 0$ and α_T^0 being the LORD critical values.

13. Concrete examples of discrete tests

In this section, we describe in detail some examples of discrete tests⁷.

 $^{^7 \}rm more$ details for running examples in R can be found in the vignette <code>https://cran.r-project.org/web/packages/DiscreteFDR/vignettes/DiscreteFDR.html</code>

13.1. Exact Poisson test

Let $X \sim \mathcal{P}(\lambda)$ with $\lambda > 0$, and consider the one-sided testing problem of the null hypothesis " $\lambda = \lambda^0$ " against the alternative " $\lambda < \lambda^0$ " for some pre-specified value $\lambda^0 > 0$. The p-value is defined as p = G(X) where $G(\cdot)$ is the c.d.f. of the distribution $\mathcal{P}(\lambda^0)$. The support $\mathcal{S} = G(\mathbb{N}) \subset [0,1]$ is finite so that p is said to be discrete and the discrete distribution function of p is defined in the usual way as $F(s) = P_{\lambda^0}(p \leq s) = \sum_{q \leq s, q \in \mathcal{S}} \mathbf{P}(p = q)$ and it can be shown that F(s) = s for $s \in [0,1] \cap \mathcal{S}$ and F(s) < s for $s \in [0,1] \setminus \mathcal{S}$.

13.2. Permutation test

Let us observe two independent samples of real random variables: X_1, \ldots, X_{n_1} i.i.d. $\sim Q_1$ and $X_{n_1+1}, \ldots, X_{n_1+n_2}$ i.i.d. $\sim Q_2$. We want to test the null " $Q_1 = Q_2$ " versus the alternative " $Q_1 \neq Q_2$ " by using some test statistic S(X) so that a large value of S(X) provides evidence against the null, for instance $S(X) = \left| n_1^{-1} \sum_{i=1}^{n_1} X_i - n_2^{-1} \sum_{i=n_1+1}^{n_1+n_2} X_i \right|$.

For any permutation σ of $\{1,\ldots,n\}$ (with $n=n_1+n_2$), we let X^{σ} denote a permutation of $X=(X_1,\ldots X_{n_1},X_{n_1+1},\ldots,X_{n_1+n_2})$ and $S(X^{\sigma})$ the corresponding test statistic. Now, a (permutation-based) p-value can be computed as follows:

$$p(X) = (B+1)^{-1} \left(1 + \sum_{b=1}^{B} \mathbf{1} \{ S(X^{\sigma_b}) \ge S(X) \} \right),$$

where $\sigma_1, \ldots, \sigma_B$ are i.i.d. permutations, independent of X, and uniformly distributed on the set of all permutations of $\{1,\ldots,n\}$, for some number $B \geq 1$ fixed in advance. Here, the support of p(X) is $S = \{1/(B+1),\ldots,1\}$ and a bounding c.d.f. of p(X) under the null is given by

$$\mathbf{P}(p(X) \le s) \le F(s) = \frac{\lfloor (B+1)s \rfloor}{B+1},$$

see Lemma 5.2 of Arlot et al. (2010).

13.3. Fisher's exact test

Fisher's exact test can be used for testing various hypotheses on count data summarized by 2×2 contingency tables (for more details, see Lehmann and Romano (2022); Agresti (2002)).

Suppose the observed 2×2 table for realizations of two Bernoulli rv X and Y is given by Table 6. Then the count n_{11} can be used as a test statistic for testing e.g. the null hypothesis H_0 that X and Y are independent. Conditioning

	Y = 1	Y = 0	Total
X = 0	n_{11}	n_{12}	n_1 .
X = 1	n_{21}	n_{22}	n_2 .
Total	$n_{\cdot 1}$	$n_{\cdot 2}$	n

on both sets of margin totals in Table 6 yields that N_{11} has a hypergeometric distribution under the null with

$$P_{H_0}(N_{11}=t) = \frac{\binom{n_1}{t}\binom{n_2}{n_1-t}}{\binom{n}{n}},$$

where the range of possible values of N_{11} is given by $\mathcal{X} = \{\max(0, n_1. + n_{\cdot 1} - n), \ldots, \min(n_1., n_{\cdot 1})\}$. Thus, if e.g. a small observed value of N_{11} provides evidence for the alternative, the (one-sided) p-value for Table 6 is given by $p = G(N_{11})$, where $G(\cdot)$ is the c.d.f. of N_{11} under the null. Thus, for fixed marginal totals, the support of the rv p is given by the finite set $\mathcal{S} = G(\mathcal{X}) \subset [0,1]$. Therefore, p is discretely distributed and the discrete distribution function of p is defined in the usual way as $F(s) = P_{H_0}(p \leq s) = \sum_{q \leq s, q \in \mathcal{S}} P(p = q)$ and it can be shown that F(s) = s for $s \in [0,1] \cap \mathcal{S}$ and F(s) < s for $s \in [0,1] \setminus \mathcal{S}$.

14. Additional numerical experiments for Section 5

14.1. Sample size

Figure 12 illustrates results when the sample size N, i.e., the subjects number per group, takes values in the set $\{25, 50, \ldots, 150\}$. As expected, the power plots show that the detection problem becomes easier when N increases. In fact, for large N the power of all procedures converge to 1. We see that our rewarded procedures do well on the whole range of N values and improve substantially on existing OMT procedures for small and moderate values of N, including our default value N=25.

14.2. Signal strength

In Figure 13, we vary the strength of the signal p_3 in the set $\{0.1, 0.2, ..., 1\}$. We see that the SUR procedures dominate their base counterparts, as expected. In addition, depending on the signal strength, the gain in power can be considerable. Also note that, perhaps surprisingly, all curves exhibit a decrease in power for p_3 near 1. Since this happens even for the original OB procedure, this is not due to the super-uniformity reward, but could perhaps be caused by the behavior of the power function of multiple Fisher exact tests taken at different levels.

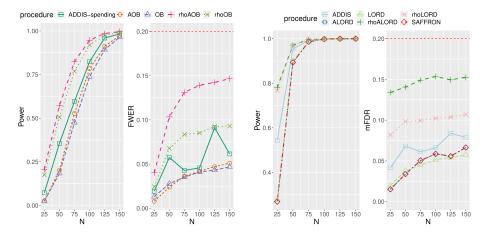


Fig 12. Power and type I error rates of the considered procedures versus $N \in \{25, 50, \dots, 150\}$, the number of subjects in the groups.

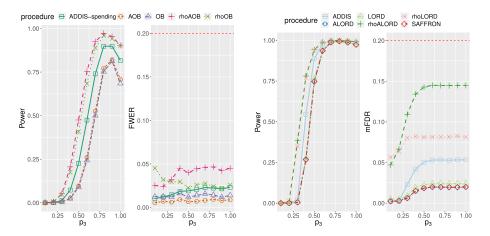


Fig 13. Power and type I error rates of the considered procedures versus the strength of the signal $p_3 \in \{0.1, 0.2, \dots, 0.9, 1\}$.

14.3. Local alternatives

As Figure 12 demonstrates, for a fixed value of the signal strength p_3 , the detection problem becomes easier as N increases, so that all procedures attain a power of 1. In this section we are interested in obtaining a more refined analysis of the various power curves when N is large. To this end, we introduce local alternatives, i.e. we now model p_3 as a function of the sample size N. To be more specific, we take $N \in \{5, 10, \ldots, 30\} \times 1000$ and set $p_3 = p_1 + \frac{1}{\sqrt{N}}$ for mFDR procedures and, $p_3 = p_1 + \frac{1.5}{\sqrt{N}}$ for FWER procedures, we fix $p_1 = p_2 = 0.1$, and generate simulated data as in Section 5.2. Figure 14 displays power and error

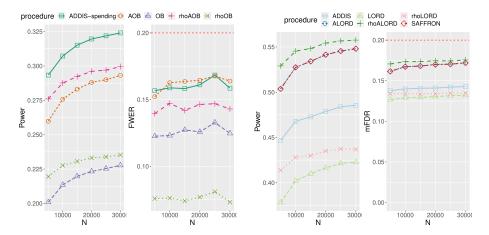


Fig 14. Power and type I error rates of the considered procedures versus $N \in \{5, 10, \dots, 30\} \times 1000$. with local alternatives.

rates for this data. Taking N as a (crude) proxy for discreteness, we observe that even with a low discreteness (say $N \leq 30000$) the SUR methods still provide some degree of improvement. Finally, for FWER procedures, ADDIS-spending provides the best power performance over the whole range of the experiment. This might be explained by the setting causing very conservative nulls p-values (i.e. very close to 1), thus allowing the discarding scheme to redistribute and spend a large part of the wealth on testing alternative hypotheses. Using the SUR method along with the discarding scheme (Tian and Ramdas, 2019, 2021) might provide an interesting avenue for further improvement, but this would define yet another class of procedures, which is outside of the scope of this paper.

14.4. Adaptivity parameter

We study the choice of λ for the adaptive procedures. In the literature, different choices of λ were proposed: in the offline case, $\lambda=0.5$ was often used as a default choice, while $\lambda=\alpha$ was recommended under dependence, see Blanchard and Roquain (2009). In the online case, Tian and Ramdas (2021) used $\lambda=\alpha$ for FWER while Ramdas et al. (2018) used $\lambda=1/2$ for mFDR. In the latter reference, the authors showed that the performance of a specific choice of λ depends on the alternative distribution in general.

Our results are presented in Figure 15 for the different adaptive procedures considered in this paper and for our particular numerical experiment setting. We can see that the SUR approach is improving the performances across all the considered values of λ , with a slightly better overall power for smaller values of λ . This experiment also shows that the default choice $\lambda = 1/2$ in our experiments does not play any specific role in the performance of our SUR pro-

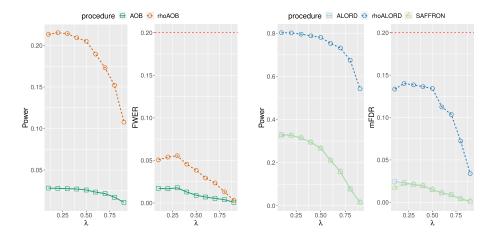


Fig 15. Power and type I error rates, for the considered procedures, versus the adaptivity parameter λ .

cedures. It is mainly motivated by previous literature to calibrate the adaptive base procedures.

14.5. Rectangular kernel bandwidth

Finally, we study the choice of the bandwidth parameter h for the rectangular kernel (9) used for the rewarded procedures, see Figure 16. As we can see, using a small bandwidth provides the best performance for the mFDR controlling procedures, whereas FWER controlling procedures require a larger bandwidth.

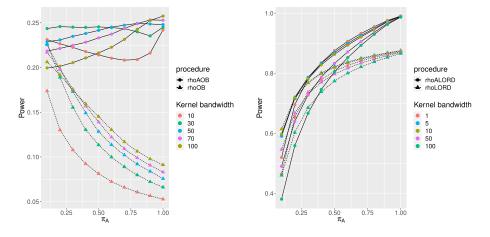


FIG 16. Power for FWER (left) and mFDR (right) rewarded procedures versus the proportion of signal π_A , for different kernel bandwidths.

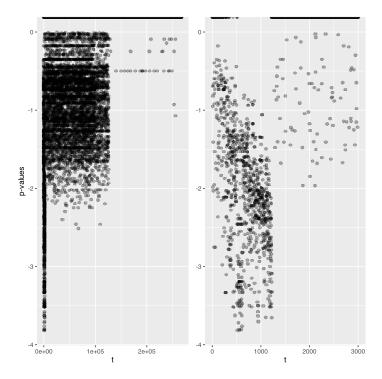


Fig 17. p-values for male mice in the IMPC data of Section 5.3. The left panel presents all p-values, the right panel the first 3000 p-values. The p-values have been transformed as in Figure 3.

The choices h=100 for FWER controlling procedures, and h=10 for mFDR controlling procedures seem reasonable although not necessarily optimal. Our experiments also show that the performance is quite sensitive to the choice of bandwidth.

15. Additional figures for the analysis of IMPC data

15.1. Localization of small p-values

Figures 17 and 18 show that small p-values mostly occur at the beginning of the data set, both for male and female mice.

15.2. Figures for female mice in the IMPC data

Figures 19 and 20 display the critical values of the studied online procedures when applied to the IMPC data in the case of female mice.

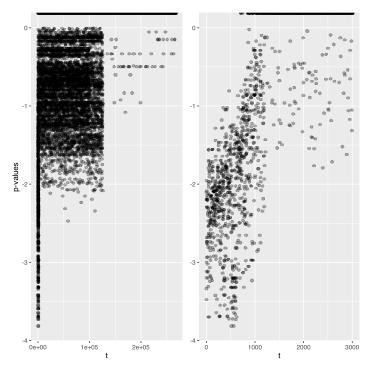


Fig 18. p-values for female mice in the IMPC data of Section 5.3. The left panel presents all p-values, the right panel the first 3000 p-values. The p-values have been transformed as in Figure 3.

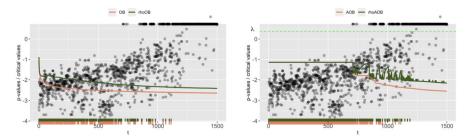


Fig 19. Same as Figure 7 but for female mice of IMPC data (see Section 5.3).

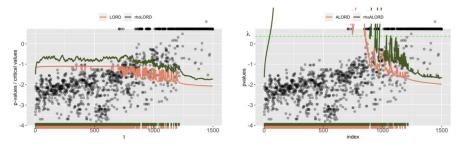


Fig 20. Same as Figure 8 for female mice of IMPC data (see Section 5.3).

16. An experiment with permutation tests

We consider the permutation test setting described in Section 13.2, with $m=1\,000$ tests, $m_0=0.6m=600$ nulls, $n_1=10,\ n_2=10,\ Q_1=\mathcal{N}(0,1)$ and $Q_2=\mathcal{N}(\mu,1)$, with $\mu=3$. The $m_1=400$ alternatives appear as follows in the stream: 300 take position $\{1,\ldots,300\}$ and then the remaining 100 are randomly scattered at positions $\{301,\ldots,500\}$. Since the reward is always smaller than 1/(B+1) by construction, we expect a significant improvement only when B is not too large, and therefore we set the number of permutations to B=20 (across all tests), and we run the procedures for $\alpha=0.5$. These choices are rather unusual but they are only used to provide a proof of concept.

While providing a more convincing case or real data application is left for future investigation, our preliminary findings presented in Figure 21 show that our SUR approach can lead to a few more rejections.

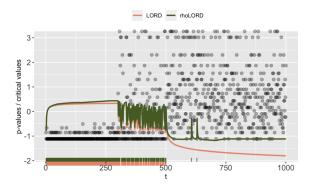


FIG 21. Same representation as in Figure 8. The procedures are run on simulated data as described in Appendices 13.2 and 16. The SUR approach makes 5 additional rejections.

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