1 1. TITLE

2 Contrast detection is enhanced by deterministic, high-frequency transcranial 3 alternating current stimulation with triangle and sine waveform.

- 4
- 5 2. ABBREVIATED TITLE

6 tACS modulates visual processing of V1.

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52 Abstract

53 Stochastic Resonance (SR) describes a phenomenon where an additive noise (stochastic 54 carrier-wave) enhances the signal transmission in a nonlinear system. In the nervous 55 system, nonlinear properties are present from the level of single ion channels all the way to 56 perception and appear to support the emergence of SR. For example, SR has been 57 repeatedly demonstrated for visual detection tasks, also by adding noise directly to cortical 58 areas via transcranial random noise stimulation (tRNS). When dealing with nonlinear 59 physical systems, it has been suggested that resonance can be induced not only by adding 60 stochastic signals (i.e., noise) but also by adding a large class of signals that are not 61 stochastic in nature which cause "deterministic amplitude resonance" (DAR). Here we 62 mathematically show that high-frequency, deterministic, periodic signals can yield 63 resonance-like effects with linear transfer and infinite signal-to-noise ratio at the output. We 64 tested this prediction empirically and investigated whether non-random, high-frequency, 65 transcranial alternating current stimulation applied to visual cortex could induce resonance-66 like effects and enhance performance of a visual detection task. We demonstrated in 28 67 participants that applying 80 Hz triangular-waves or sine-waves with tACS reduced visual 68 contrast detection threshold for optimal brain stimulation intensities. The influence of tACS 69 on contrast sensitivity was equally effective to tRNS-induced modulation, demonstrating that 70 both tACS and tRNS can reduce contrast detection thresholds. Our findings suggest that a 71 resonance-like mechanism can also emerge when deterministic electrical waveforms are 72 applied via tACS.

73 Keywords

deterministic amplitude resonance; stochastic resonance; high-frequency transcranial
 alternating current stimulation; neuromodulation; visual processing; contrast sensitivity;
 transcranial random noise stimulation

77 New & Noteworthy

Our findings extend our understanding of neuromodulation induced by noninvasive electrical stimulation. We provide first evidence showing acute online benefits of tACS_{triangle} and tACS_{sine} targeting the primary visual cortex (V1) on visual contrast detection in accordance with the resonance-like phenomenon. The 'deterministic' tACS and 'stochastic' hf-tRNS are equally effective in enhancing visual contrast detection.

83 **1. Introduction**

84 **1.1 On stochastic resonance**

85 Stochastic resonance (SR) was discovered in the context of the hysteresis features of 86 climate (ice age) (1-3). Since then it has been generalized and studied in a variety of 87 naturally occurring processes including biological systems (4, 5). Demonstrations of SR in 88 the nervous system were carried out on crayfish mechanoreceptors (6), neurons in crickets 89 (7), mice (8, 9), rats (10–12), cats (13), and humans (see 1.3 Stochastic resonance effects 90 on neural processing below) with studies consistently reporting enhanced system 91 performance. Signal enhancement is described in a vast body of literature as the basic 92 property of resonance mechanisms (14). Here we survey a few basic features of SR that are 93 directly relevant for our paper. In general, the quality of signal transfer through a system is 94 characterized by the following parameters at the output: amplification (or the signal strength), 95 linearity, signal-to-noise ratio, and the phase shift.

96 SR is a phenomenon where the transfer of a periodic or aperiodic signal in a nonlinear 97 system is optimized by an additive -typically Gaussian- noise (15). Note that originally, when 98 SR was studied in binary systems, it represented a frequency-resonance, that is, matching 99 the period time of the periodic signal with the mean residence time in the potential wells of 100 the binary system driven by a stochastic carrier-wave (noise). Later the argument behind the 101 name SR was modified to amplitude-resonance. Today, "resonance" means an optimal root-102 mean-square (RMS) amplitude value of the noise, i.e., amplitude-resonance at the carrier-103 wave RMS amplitude level for the best signal transmission.

104 In the initial phase of SR research, the nonlinear systems were bistable (1). At a later stage it 105 was discovered that monostable systems (including neurons) also offer SR (16). Moreover, it 106 was realized that the memory/hysteresis effects of the bistable systems actually cause a 107 stochastic phase shift (phase noise) that negatively impacts the quality of the transferred 108 signal (17). Due to this fact, the best stochastic resonators are the memory-free Threshold 109 Elements (TE), such as the Level Crossing Detector (LCD) (18) and the Comparator (19). 110 The LCD device (the simplest model of a neuron) produces a short, uniform spike whenever 111 its input voltage amplitude is crossing a given threshold level in a chosen, typically positive 112 direction. On the other hand, the Comparator has a steady binary output where the actual 113 value is dictated by the situation of the input voltage amplitude compared to a given 114 threshold level: for example, in the sub-threshold case the output is "high" while in the supra-115 threshold case, it is "low".

116 At the output of a stochastic resonator, the signal strength (SS), the signal-to-noise-ratio 117 (SNR), the information entropy and the Shannon information channel capacity show maxima 118 versus the intensity of the additive input noise. However, these maxima are typically located 119 at different noise intensities. Exceptions are the SNR and information entropy which are 120 interrelated by a monotonic function; thus they have the same location of their maxima, see 121 the arguments relevant for neural spike trains (20). On the other hand, the information 122 channel capacity of SR in an LCD and in neural spike trains has the bandwidth as an extra 123 variable controlled by the input (the higher the input noise the higher the bandwidth); thus 124 the different location of its maximum is at higher input noise than for the maximum of the 125 SNR (21).

126 It is important to note that, in the linear response limit, that is, when the input signal is much 127 smaller than the RMS amplitude of the additive carrier-wave (Gaussian noise), the SNR at 128 the output is always less than at the input (see the mathematical proof in (15)). 129 Consequently, the information content at the output is always less than at the input. On the 130 contrary, in the nonlinear response limit, the SNR at the output can be enhanced by several 131 orders of magnitude compared to its input value provided the signal has a small duty cycle, 132 such as neural spikes do (17, 22). Yet, due to the unavoidable noise at the output, which is 133 the unavoidable impact of the stochastic carrier-wave (noise), the information at the output (signal plus noise) is always less than in the *original* input signal *without the added carrier- wave* (*noise*).

136 Therefore, if a proper additive, high-frequency, periodic time function could be used as 137 carrier-wave in a stochastic resonator instead of a Gaussian noise, the fidelity and the 138 information content of the input signal could be preserved while it is passing through the 139 nonlinear device, as we will show below. However, even in this case there is an optimal 140 (range) for the mean-square amplitude of the carrier-wave. Thus, we call this deterministic 141 phenomenon "deterministic amplitude resonance", (DAR), which is also an amplitude-142 resonance where a deterministic (instead of stochastic/noise) carrier-wave with sufficiently 143 large amplitude produces the optimal signal transfer via the system.

144 **1.2** Deterministic amplitude resonance (DAR) with high-frequency periodic carrier 145 waves

First Landa and McClintock (23) realized that SR like phenomena could occur with highfrequency sinusoidal signals instead of noise. They successfully demonstrated their idea by computer simulations of a binary (double-well potential) SR system. Recently, Mori, et al (24) used high-frequency, noise-free, periodic neural spikes for excitation in a neural computer model to show that SR like features on the mutual information can be achieved by tuning the frequency of these periodic excitation in the 80-120 Hz range.

Below, we show that high-frequency triangle waves can offer a noise-free signal transfer which can be exactly linear at certain conditions. Sinusoidal waves are also discussed briefly.

155 **1.2.1** The case of triangle (or sawtooth) carrier-waves, instead of noise

Earlier, in a public debate about the future of SR, one of us proposed a noise-free method byutilizing high-frequency triangle waves to improve signal transmission through threshold

devices (25) and to reach exactly linear transfer and infinite SNR at the output. Here wesummarize those arguments.

Figure 1 shows an example of stochastic resonator hardware with an additive triangle wave, as the carrier-wave, instead of noise. The same argumentation works for sawtooth wave, too. Note: the original threshold-based stochastic resonators (17, 18) contain the same hardware elements where Gaussian random noise is used instead of the triangle wave. Due to the binary nature of the visual detection experiments described in this paper our focus is on sub-threshold binary (square-wave) signals with some additional comments about the case of analog signals.

167

FIGURE 1

168 The TE is either an LCD or a Comparator. Suppose that the stable output of the LCD is zero 169 and it produces a short uniform positive spike with height $U_{
m LCD}$ and duration au whenever 170 the input level crosses the Threshold in upward direction. The Comparator's output stays at 171 a fixed positive value whenever the input level is greater than the threshold and stays at a 172 lower value (zero or negative) otherwise. Suppose when the input level is greater than the 173 Threshold, $U_{\rm th}$, the Comparator output voltage $U_{\rm c} = U_{\rm H}$ and otherwise it is 0. The Low-pass 174 Filter creates a short-time moving-average in order to smooth out the high-frequency 175 components (frequency components due to switching triggered by the carrier wave) and it 176 keeps only the low-frequency part which is the bandwidth of the signal. The parameters, 177 such as the frequency f_s of the signal, the frequency f_t of the triangle wave and the cut-off frequency f_{c} of the Low-pass Filter should satisfy 178

179
$$f_{\rm s} << f_{\rm c} << f_{\rm t} < \frac{1}{\tau}$$
 (1)

180 in order to transfer the signal with the highest fidelity.

181 The upper part of **Figure 1** shows the situation without carrier wave: the sub-threshold 182 binary signal is unable to trigger the TE thus the output signal is steadily zero. The lower part 183 of Figure 1 shows the situations where an additive, triangle wave assists the signal to reach 184 the threshold thus it carries the binary signal over the TE resulting in a nonzero output 185 signal. The triangular wave will have to be of a high enough frequency for two main reasons 186 i) because of the Nyquist sampling theorem, the sampling frequency needs to be at least 187 twice as large than the highest frequency component of the signal, ii) for the low-pass filter to 188 be able to smooth out the carrier signal (which is a trash), the carrier signal frequency must 189 be much larger than the reciprocal of the time duration of the binary signal.

i) The case of Level Crossing Detector (LCD)

191 If a constant input signal plus triangle wave can cross the threshold, the LCD produces a 192 periodic spike sequence with the frequency of the triangle wave. In this situation, the time 193 average of this sequence is $f_t \tau U_{LCD}$ therefore, for the binary input signal, the output of the 194 LPF will be binary with amplitude values:

195
$$U_{\text{LPF}}(t) = f_t \tau U_{\text{LCD}} \quad \text{or} \quad 0 \tag{2}$$

Thus, the binary input signal is restored at the output of the LPF without any stochasticity (noise) in it. The only deviation from the input signal is a potentially different amplitude (nonzero amplification) and some softening of the edges dues to the LPF depending on how well Relation 1 is satisfied.

200 In conclusion, with an LCD as TE, regarding the amplitude resonance versus the carrier 201 wave amplitude U_t , there are three different input amplitude ranges:

202 (a) $U_{s} + U_{t} < U_{th}$ then there is no output signal

203 (b) $U_{th} < U_s + U_t$, $U_s < U_{th}$, $U_t < U_{th}$ then the binary signal is restored at the output

204 (c)
$$U_{\rm th} < U_{\rm t}$$
 then the output is steadily at the high level $U_{\rm LPF}(t) = f_{\rm t} \tau U_{\rm LCD}$

Therefore, the binary signal can propagate to the output only in the (b) situation when it does that without any noise contribution at the output (the SNR is infinite).

207 ii) The case of Comparator

Note, this system is very different from "Stocks's suprathreshold SR" (19), where a large number of independent comparators with independent noises are used with a common signal and an adder to reach a finite SNR. For the sake of simplicity, but without limiting the generality of the argumentation, suppose that the binary signal, $U_s(t)$, values are 0 and U_s , where $U_s \leq U_{th}$, and the maximum amplitude of the triangle signal, $U_t(t)$ is U_t and its minimum value is 0. In conclusion:

214 when
$$U_{\rm s}(t) + U_{\rm t}(t) > U_{\rm th}$$
, $U_{\rm c} = U_{\rm H}$ otherwise $U_{\rm c} = 0$ (3)

i.e., the comparator's output voltage has only 2 possible steady states: 0 and U_{H} . When the input voltage is above the threshold voltage U_{th} the output is U_{H} , otherwise it is 0.

To evaluate the average output voltage of the comparator the first question is the fraction of time that the input spends over the threshold, see **Figure 2**.

219 ### FIGURE 2 ###

This time $T_{\rm H}$ within a period of the triangle wave is the period duration $1/f_{\rm t}$ minus twice the time $t_{\rm r}$ spent for rising from the minimum to the threshold:

222
$$T_{\rm H} = \frac{1}{f_{\rm t}} - 2 t_{\rm r} = \frac{1}{f_{\rm t}} - 2 \frac{U_{\rm th} - U_{\rm s}}{2U_{\rm t} f_{\rm t}} = \frac{U_{\rm t} - U_{\rm th} - U_{\rm s}}{U_{\rm t} f_{\rm t}},$$
(4)

223 where we used that the slope , of the triangle signal with peak-to-peak amplitude is

$$s = 2f_t U_t \quad , \tag{5}$$

assumed that the signal amplitude U_s is present at the input and assumed condition (3) that the signal alone is subthreshold, but the sum of the signal and the triangle wave is suprathreshold:

From (3) and (4), the smoothed value of the output voltage $U_{LPF}(t)$ of the LPF when the input signal amplitude is :

231
$$U_{\rm LPF} = \langle U_{\rm LPF}(t) \rangle = U_{\rm H} \frac{T_{\rm H}}{1/f_{\rm t}} = U_{\rm H} \frac{U_{\rm t} - U_{\rm th} + U_{\rm s}}{U_{\rm t}} = U_{\rm H} \frac{U_{\rm t} - U_{\rm th}}{U_{\rm t}} + \frac{U_{\rm H}}{U_{\rm t}} U_{\rm s} , \qquad (7)$$

232 where denotes short-range averaged (smoothed) value discussed above.

The last term in the right side of Equation (7) demonstrates that the signal amplitude transfers linearly through the system. Therefore, this version of our device is working distortion-free also for analog signals, not only for the present digital signal assumption.

Thus, this device is not only noise-free but also ideally linear for subthreshold signals satisfying condition (6), even though exact linearity is not an important feature during the experimental study in the present paper.

In conclusion, with a comparator as TE, regarding the amplitude resonance versus thecarrier wave amplitude , there are two different input amplitude ranges:

241 (a) $U_{\rm s} + U_{\rm t} < U_{\rm th}$ then there is no output signal

(b) $U_{th} < U_s + U_t$, $U_s < U_{th}$, then the binary signal is restored at the output and its amplitude scales inversely with the amplitude U_t of the carrier wave. The maximal amplitude is at .

Therefore, the binary signal can propagate to the output only in the (b) situation when it does that without any noise (the SNR is infinite) and it has a linear transfer for analog signals. Of note, for a large U_t (trianguler waveform) the binary signal is lost as the output would be a constant signal.

248

249 **1.2.2** The case of sinusoidal carrier waves, instead of triangle waves

250 The above argumentations qualitatively work also for sinusoidal carrier-waves except that 251 the linearity of the transfer is lost. The triangle carrier-wave has a Fourier series that has only odd harmonics, where the *n*-th harmonic amplitudes scale with $1/n^2$, that is, the 252 253 strongest harmonic (the 3-rd) is 9 times weaker, and the next strongest harmonic (the 5-th) 254 is 25 times less than the base harmonic. The qualitative difference is that the absolute value 255 of the slope of sinusoidal carrier-wave is reduced when approaching its peak level and it is 256 zero at the peak. The constant slope of the triangle wave is essential for the exactly linear 257 transfer, see the mathematical proof above.

In conclusion, when sinusoidal carrier-wave is used instead of a triangle (or sawtooth) wave, the same qualitative features remain, including the zero-noise contribution at the output (infinite SNR). The exception is the linearity of transfer of analog signals via the comparator which is lost at sinusoidal carrier-wave.

262 1.3 Stochastic resonance effects on neural processing

In neural systems, it has been demonstrated that responses to externally applied stimuli were maximally enhanced when an optimal level of electrical random noise stimulation was applied. These effects were linked specifically to the opening of voltage gated sodium (Na⁺) channels in response electrical stimulation, causing a sodium influx, which in turn causes alocal depolarization of the cell membrane (9, 12, 26).

In humans, early SR effects have been mainly demonstrated via behavioral signal detection
tasks whereby noise was added to the periphery. For example, the detection of low-contrast
visual stimuli was significantly enhanced when the stimuli were superimposed with visual
noise (27)

272 Recently, similar enhancements of visual perception have been reported when noise was 273 directly added to the cerebral cortex by the means of transcranial random noise stimulation 274 (tRNS) in studies investigating its acute effects on visual processing (26, 28-33). According 275 to the SR theory, while the optimal level of tRNS benefits performance, excessive noise is 276 detrimental for signal processing (28, 29, 33), resulting in an inverted U-shape relationship 277 between noise benefits and noise intensity. In consistence with SR, tRNS was shown to be 278 particularly beneficial for visual detection performance when the visual stimuli were 279 presented with near-threshold intensity (28, 29, 34).

280 However, based on the theoretical consideration described above, a resonance-like 281 phenomenon can be observed with deterministic stimulations. Here we test this prediction 282 empirically and investigate if the response of visual cortex to around-threshold contrast 283 stimuli could also be enhanced via a high-frequency deterministic signal. We tested if 284 triangle or sine waves can modulate signal processing in a resonance-like manner by 285 delivering tACS with triangle waveform (tACS_{triangle}) or sine waveform (tACS_{sine}) targeting the 286 primary visual cortex (V1) of participants performing a visual contrast sensitivity task and 287 measured their visual detection threshold. We hypothesized that resonance-like DAR effects 288 would be reflected in the beneficial influence of high-frequency stimulation on signal 289 processing via signal enhancement.

290 **2. Materials and methods**

12

291 2.1 Participants

292 Individuals with normal or corrected-to-normal vision and without identified contraindications 293 for participation according to established brain stimulation exclusion criteria (35, 36) were 294 recruited in the study. All study participants provided written informed consent before the 295 beginning of each experimental session. Upon study conclusion participants were debriefed 296 and financially compensated for their time and effort. All research procedures were approved 297 by the Cantonal Ethics Committee Zurich (BASEC Nr. 2018-01078) and were performed in 298 accordance with the Helsinki Declaration of the World Medical Association (2013 WMA 299 Declaration of Helsinki) and guidelines for non-invasive brain stimulation research through 300 the COVID-19 pandemic (37).

301 The required sample size was estimated using an a priori power analysis (G*Power version 302 3.1; (38)) based on the effect of maximum contrast sensitivity improvement with tRNS shown by Potok et al. (39) (η_p^2 = 0.165, Effect size f = 0.445). It revealed that 28 participants should 303 304 be included in an experiment to detect an effect with repeated measures analysis of variance 305 (rmANOVA, 4 levels of stimulation condition), alpha = 0.05, and 90% power. We included 31 306 participants in experiment 1 (tACS_{triangle}) and 32 participants in experiment 2 (tACS_{sine}) to 307 account for potential dropouts. Visual contrast detection is potentially prone to floor effects if 308 the contrast detected at baseline approaches the technical limits of the setup. We decided to 309 exclude participants that are exceptionally good in the visual task and present visual contract 310 threshold below 0.1 (Michelson contrast, see Visual stimuli) in the baseline, no tACS 311 condition (for visual contrast intensity range of minimum 0 and maximum 1) in the no tACS 312 baseline condition. For outliers' removal we used standardized interquartile range (IQR) 313 exclusion criteria (values below Q1-1.5IQR or above Q3+1.5IQR, where Q1 and Q3 are 314 equal to the first and third quartiles, respectively) to avoid accidental results, unlikely driven 315 by tES, e.g., due to participants responding without paying attention to the task.

From the initially recruited sample, we excluded 7 individuals. In tACS_{triangle} experiment 1: 1 participant revealed exceptional contrast threshold modulation (>Q3+1.5IQR), 1 participant had a contrast threshold below 0.1 in the baseline condition (also >Q3+1.5IQR), 1 participant stopped the session because of unpleasant skin sensations. In tACS_{sine} experiment 2: 1 participant revealed exceptional contrast threshold modulation (>Q3+1.5IQR), 1 participant stopped the session because of unpleasant skin sensations. In tACS_{sine} experiment 2: 1 participant revealed exceptional contrast threshold modulation (>Q3+1.5IQR), 1 participant stopped the session because of unpleasant skin sensations, 2 participants reported frequent (75% accuracy) phosphenes sensation due to stimulation (see *tACS characteristics*).

The final sample consisted of 28 healthy volunteers (16 females, 12 males; 26.9 \pm 4.7, age range: 21-39) in tACS_{triangle} experiment 1, and 28 healthy volunteers (20 females, 8 males; 26.4 \pm 4.4, age range: 20-39) in tACS_{sine} experiment 2. We did not collect information about the race of participants. Twenty of these participants completed both experimental sessions. For participants who took part in both experiments, 15 participants started with tACS_{triangle} and 5 with tACS_{sine}. The experimental sessions took place on different days with 2.6 \pm 1.2 months on average apart. Delays were caused by COVID-19 pandemic (37).

330 2.2 General Study design

331 To evaluate the influence of tACS on visual contrast detection, we performed two 332 experiments in which we delivered either tACS_{triangle}, or tACS_{sine} targeting V1, during visual 333 task performance (see **Figure 3A**). In each experiment, three tACS intensities and a control 334 no tACS condition were interleaved in a random order. Our main outcome parameter in all 335 experiments was a threshold of visual contrast detection (VCT) that was determined for each 336 of the different tACS conditions (39). The experimental procedure to estimate VCT followed 337 a previously used protocol to assess the influence of tRNS on contrast sensitivity (39). In 338 brief, VCT was estimated twice independently, in two separate blocks within each session 339 (see Figure 3D). We determined the individual's optimal tACS intensity (defined as the 340 intensity causing the lowest VCT, i.e., biggest improvement in contrast sensitivity) for each 341 participant in the 1st block of experiment 1 (ind-tACS_{triangle}) and experiment 2 (ind-tACS_{sine}) and retested their effects within the same experimental session on VCT data acquired in the
 2nd block.

344

FIGURE 3

345 2.2.1 Experimental setup and visual stimuli

346 The experiments took place in a dark and quiet room, ensuring similar lighting conditions for 347 all participants. Participants sat comfortably, 0.85m away from a screen, with their head 348 supported by a chinrest. Visual stimuli were generated with Matlab (Matlab 2020a, 349 MathWorks, Inc., Natick, USA) using the Psychophysics Toolbox extension that defines the 350 stimulus intensity with Michelson contrast (40-42) and displayed on a CRT computer screen 351 (Sony CPD-G420). The screen was characterized by a resolution of 1280 x 1024 pixels, refresh rate of 85Hz, linearized contrast, and a luminance of 35 cd/m² (measured with J17 352 353 LumaColor Photometer, TektronixTM). The target visual stimuli were presented in the form of 354 the Gabor patch – a pattern of sinusoidal luminance grating displayed within a Gaussian 355 envelope (full width at half maximum of 2.8 cm, i.e., 1° 53' visual angle, with 7.3 cm, i.e., 4° 356 55' presentation radius from the fixation cross, staying within the central vision, i.e., <8° 357 radius; (43, 44)). The Gabor patch pattern consisted of 16 cycles with one cycle made up of 358 one white and one black bar (grating spatial frequency of 8 c/deg). Stimuli were oriented at 359 45° tilted to the left from the vertical axis (see **Figure 3A**), since it was shown that tRNS 360 enhances detection of low contrast Gabor patches especially for non-vertical stimuli of high 361 spatial frequency (31).

362 **2.2.2** Four-alternative forced choice visual detection task

In both experiments, participants performed a visual four-alternative forced choice (4-AFC)
visual task, designed to assess an individual VCT, separately for each stimulation condition.
A 4-AFC protocol was shown to be more efficient for threshold estimation than commonly
used 2-AFC (45). Participants were instructed to fixate their gaze on a cross in the center of

367 the screen. In the middle of each 2.04s trial, a Gabor patch was randomly presented for 368 40ms in one of the 8 locations (see Figure 3A). A stimulus appeared in each location for the 369 same number of times (20) within each experimental block in pseudo-randomized order to 370 avoid a spatial detection bias. The possible locations were set on noncardinal axes, as the 371 detection performance for stimuli presented in this way is less affected (i.e. less variable) 372 than when stimuli are positioned on the cardinal axes (46). Each trial was followed by 1s 373 presentation of only fixation cross after which the 'response screen' appeared. Participants' 374 task was to decide in which quadrant of the screen the visual stimulus appeared and indicate 375 its location on a keyboard (see Figure 3A). The timing of the response period was self-376 paced and not limited. Participants completed a short training session (10 trials), with the 377 stimulus presented always at high contrast (0.5; for visual contrast intensity range of 378 minimum 0 and maximum 1), in order to ensure that they understand the task (see Figure 379 3D).

380 VCT was estimated using the QUEST staircase procedure (47), implemented in the 381 Psychophysics Toolbox in Matlab (40-42), which is a method used in psychophysical 382 research to estimate threshold of a psychometric function (47). The thresholding procedure 383 starts with a presentation of the visual stimulus displayed with 0.5 contrast intensity 384 (Michelson contrast, for visual contrast intensity ranging 0-1; note that the stimuli were 385 displayed for just 40ms). When participants answer correctly, QUEST lowers the presented 386 contrast intensity. Consequently, when participants answer incorrectly QUEST increases the 387 presented contrast. The estimated contrast intensity for the next stimulus presentation is 388 based on a maximum-likelihood-based estimate of the underlying psychometric function. 389 Characteristics of the stimuli on each trial are determined by the input stimuli and respective 390 system responses that occurred in the previous sequence of trials (48). The estimated 391 stimulus contrast is adjusted to yield 50% detection accuracy (i.e., detection threshold 392 criterion, see Figure 3C). For a 4-AFC task 25% accuracy corresponds to a chance level. 393 The remaining parameters used in the QUEST staircase procedure where set as follows: 394 steepness of the psychometric function, beta = 3; fraction of trials on which the observer 395 presses blindly, delta = 0.01; chance level of response, gamma = 0.25; step size of internal 396 table grain = 0.001; intensity difference between the largest and smallest stimulus intensity, 397 range = 1. VCT was assessed across 40 trials per stimulation condition. Four different 398 conditions were randomly interleaved within each of 2 experimental blocks (40 trials x 4 399 conditions x 2 blocks; total number of 320 trials per experimental session, **Figure 3D**).

400 2.2.3 tACS characteristics

401 In stimulation trials, tACS (80Hz) with symmetrical triangle- (tACS_{triangle}) or sinewave 402 (tACS_{sine}), with no offset was delivered. Stimulation started 20ms after trial onset and was 403 maintained for 2s (see Figure 3A). Subsequently a screen with only fixation cross was 404 displayed for 1 s, followed by the self-paced response time. tACS waveforms were created 405 within Matlab (Matlab 2020a, MathWorks, Inc., Natick, USA) and sent to a battery-driven 406 electrical stimulator (DC-Stimulator PLUS, NeuroConn GmbH, Ilmenau, Germany), operated 407 in REMOTE mode, via a National Instruments I/O device USB-6343 X series, National 408 Instruments, USA). The active tACS conditions and no tACS control condition were 409 interleaved and presented in random order. Timing of the stimuli presentation, remote 410 control of the tACS stimulator, and behavioral data recording were synchronized via Matlab 411 (Matlab 2020a, MathWorks, Inc., Natick, USA) installed on a PC (HP EliteDesk 800 G1) 412 running Windows (Windows 7, Microsoft, USA) as an operating system.

In both experiments tACS (80Hz) stimulation (tACS_{triangle} in experiment 1 or tACS_{sine} in experiment 2) was delivered with 0.75mA, 1mA, and 1.5mA amplitude (peak-to-baseline), resulting in maximum current density of $60 \frac{\mu A}{cm^2}$, which is below the safety limits of $167 \frac{\mu A}{cm^2}$ for transcranial electrical stimulation (49). These intensities were selected based on previous studies investigating effects of tRNS on contrast sensitivity (28, 39).

17

Prior to electrode placement, an anesthetic cream (Emla® 5%, Aspen Pharma Schweiz GmbH, Baar, Switzerland) was applied to the intended electrodes position on the scalp to numb potential tACS-induced cutaneous sensations and diminish transcutaneous effects of stimulation. To ensure that the cream got properly absorbed, it was left on the scalp for 20 min (50, 51) during which participants completed task training (see *Four-alternative forced choice visual detection task* and **Figure 3D**).

424 To target V1 we used an electrode montage that was previously shown to be suitable for 425 visual cortex stimulation (28, 39, 52). The electrodes were placed on the head at least 20 426 min after the application of an anesthetic cream. One tACS 5x5cm rubber electrode was 427 placed over the occipital region (3 cm above inion, Oz in the 10-20 EEG system) and one 428 5x7cm rubber electrode over the vertex (Cz in the 10-20 EEG system). Electroconductive gel 429 was applied to the contact side of the rubber electrodes (NeuroConn GmbH, Ilmenau, 430 Germany) to reduce skin impedance. The impedance between the electrodes was monitored 431 and kept below 15 k Ω . We used electric field modelling to verify that our electrodes target 432 V1. Simulations were run in SimNIBS 2.1 (53) using the average MNI brain template (see 433 Figure 3B). Note, that the software enables finite-element modelling of electric field 434 distribution of direct current stimulation without taking into account the temporal 435 characteristics of the alternating current.

436 Since we used a very brief stimulation time (2 s only), fade in/out periods were not possible 437 (54). Accordingly, some participants were able to distinguish the stimulation conditions (see 438 Results). We accounted for this possible bias using a control measure and analysis of the 439 potential transcutaneous sensations. In each session, before the start of the main 440 experiment, participants were familiarized with tACS and we assessed the detectability of 441 potential cutaneous sensations (Figure 3D). The detection task consisted of 20 trials. 442 Participants received either 2s tACS (0.75, 1, and 1.5mA tACS_{triangle} in experiment 1 or 443 tACS_{sine} in experiment 2) or no tACS, to test whether they can distinguish between

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444 stimulation vs no stimulation. The task after each trial was to indicate on a keyboard whether 445 they felt a sensation underneath the tACS electrodes. In experiment 2 an additional control 446 measurement was added to assess the potential phosphene induction by the tACS 447 waveform. tACS (in lower frequency range) was previously suggested to induce visual 448 phosphenes (55, 56). The protocol was the same with the only difference that this time after 449 each trial participants indicated on a keyboard whether they perceived any visual sensations 450 while looking on the black computer screen. The determined detection accuracy (hit rates, 451 HR, defined as the proportion of trials in which a stimulation is present and the participant 452 correctly responds to it) of the cutaneous sensation (experiment 1 and 2) and phosphenes 453 (experiment 2) induced by tACS served as a control to estimate whether any unspecific 454 effects of the stimulation might have confounded the experimental outcomes (54). In the 455 control analysis we used the HRs for detection tACS stimulation conditions (separately for 456 tACS_{triangle} and tACS_{sine}) as covariates (see Statistical Analysis).

457 2.3 Statistical analysis

458 Statistical analyses were performed in IBM SPSS Statistics version 26.0 (IBM Corp.) and 459 JASP (0.16.3) unless otherwise stated. All data was tested for normal distribution using Shapiro-Wilks test of normality. Partial eta-squared (small $\eta_p^2 = 0.01$, medium $\eta_p^2 = 0.06$, 460 large η_p^2 = 0.14; (57)) or Cohen's d (small d=0.20–0.49, medium d=0.50–0.80, large d > 461 462 0.80; (58)) values are reported as a measure of effect-sizes. Standard statistics for simple 463 effects were complemented with their Bayesian equivalents using the Bayes factor (BF_{01}) 464 with $BF_{01} > 1$ indicating evidence in favor of the null hypothesis over the alternative 465 hypothesis. BF₀₁ were primarily provided to statistically confirm the lack of an effect 466 throughout the analyses. Variance is reported as SD in the main text and as SE in the 467 figures. Statistical analysis of tACS_{triangle} and tACS_{sine} effects was analogous to the one 468 performed to test hf-tRNS effects (39).

469 2.3.1 Analysis of VCT modulation in tACS_{triangle} and tACS_{sine} experiments

470 First, we tested whether baseline VCT in the no tACS condition differed across the two 471 experimental sessions using a Bayesian independent samples t-test (average baseline VCT 472 in blocks 1-2 in experiments 1-2) using the BF_{01} .

473 For all repeated measures analysis of variance (rmANOVA) models, sphericity was 474 assessed with Mauchly's sphericity test. The threshold for statistical significance was set at 475 $\alpha = 0.05$. Bonferroni correction for multiple comparisons was applied where appropriate (i.e., 476 post hoc tests; preplanned comparisons of stimulation 0.75mA, 1mA and 1.5mA vs no tACS 477 baseline).

478 To test the influence of tACS_{triangle} on contrast sensitivity, VCT data collected in experiment 1 479 (tACS_{triangle}) were analyzed with a rmANOVA with the factors of tACS_{triangle} (no, 0.75mA, 480 1mA, and 1.5mA tACS_{triangle}) and *block* (1st, 2nd). For each individual and each block, we 481 determined the maximal behavioral improvement, i.e., lowest VCT measured when 482 tACS_{triangle} was applied, and the associated "optimal" individual tACS_{triangle} intensity (ind-483 tACS_{triangle}). Note, that the ind-tACS_{triangle} was always selected from active stimulation 484 conditions (i.e., even if participants performed better in the no tACS baseline, the ind-tACS 485 intensity was defined based on the lowest VCT during stimulation). The maximal behavioral improvements in the 1st and the 2nd block were compared using a t-test (2-tailed) for 486 487 dependent measurements. Importantly, we determined ind-tACS_{triangle} in the 1st block, and then used the VCT data of the separate 2nd block to test whether the associated VCT is 488 489 lower compared to the no tACS condition using t-tests for dependent measures. Since we 490 had the directional hypothesis that VCT is lower for the ind-tACS_{triangle} intensity compared to 491 no tACS this test was 1-tailed. Determining ind-tACS_{triangle} and testing its effect on VCT in 492 two separate datasets is important to not overestimate the effect of tACS_{triangle} on visual 493 detection behavior.

Similarly, VCT data collected in experiment 2 (tACS_{sine}) was analyzed with a rmANOVA with the factor of *tACS_{sine}* (no, 0.75mA, 1mA, and 1.5mA tACS_{sine}) and the factor *block* (1st, 2nd). 496 Again, for each individual and each block, we determined the maximal behavioral 497 improvement and the associated ind-tACS_{sine}. We compared results obtained in the first and 498 second block using the same statistical tests as for the experiment 1. The maximal 499 behavioral improvements were compared using a t-test (2-tailed) for dependent 500 measurements. We examined whether the ind-tACS_{sine} determined based on the best behavioral performance in 1st block, caused VCT to be lower compared to the no tACS 501 condition when retested on the independent dataset (2nd block) using t-tests (1-tailed) for 502 503 dependent measures.

504 In both experiments to assess a general modulation of VCT induced by tACS we calculated 505 the mean change in VCT in all active tACS conditions from 1st and 2nd blocks normalized to 506 baseline no tACS condition (tACS-induced modulation).

507 To control for any potential unspecific effects of tACS we repeated the main analyses of VCT 508 (i.e., rmANOVA) with adding HRs of cutaneous sensation for all current levels (experiment 1, 509 tACS_{triangle} and 2, tACS_{sine}) and phosphene detection (experiment 2, tACS_{sine}) as covariate. 510 We also tested correlations between the average HR of cutaneous sensation (experiment 1 511 and 2) and phosphene (experiment 2) detection and average tACS-induced modulation 512 using a Pearson correlation coefficient.

513 2.3.2 Comparison of stimulation-induced VCT modulation in tACS_{triangle}, tACS_{sine}, and 514 hf-tRNS experiments

515 We compared the effects of deterministic transcranial electrical stimulation (tES, i.e., 516 tACS_{triangle} and tACS_{sine}) and stochastic tES (i.e., hf-tRNS) on VCT. The data demonstrating 517 the effect of hf-tRNS on VCT were taken from a previous study investigating the effects of hf-518 tRNS using the same behavioral paradigm (39). 519 First, we tested whether baseline VCT in the no tES (no tACS, no hf-tRNS) conditions 520 differed across the three experiments using a Bayesian independent samples t-test (average 521 baseline VCT in blocks 1-2 in tACS_{triangle}, tACS_{sine} and hf-tRNS) using the BF₀₁.

522 Next, we tested whether a general tES-induced modulation of VCT (mean of all active 523 stimulation conditions from two blocks normalized to baseline no stimulation condition) 524 differed across the three experiments using a Bayesian ANOVA (tES-induced modulation in 525 tACS_{triangle}, tACS_{sine} and hf-tRNS experiments) using the BF₀₁.

526 Finally, we depicted tES-induced modulation of VCT as paired Cohen's d bootstrapped 527 sampling distributions employing an online tool (https://www.estimationstats.com; (59)). For 528 each pair of control no tES (i.e., no tACS in tACS_{triangle}, tACS_{sine} and no hf-tRNS) and tES 529 conditions (tACS_{triangle}, tACS_{sine}, hf-tRNS) a two-sided permutation t-tests were conducted. 530 5000 bootstrap samples were taken. The confidence interval was bias-corrected and 531 accelerated. The reported P values are the likelihoods of observing the effect sizes, if the 532 null hypothesis of zero difference is true. For each permutation P value, 5000 reshuffles of 533 the control and test labels were performed.

534 3. Results

We first tested whether VCT measured during the no tACS conditions differed between the experiments (i.e., average baseline VCT in tACS_{triangle} and tACS_{sine} experiments, see **Figure 4**). Bayesian independent samples t-test revealed that the baseline VCT measured in the no tACS condition did not differ between experiments ($BF_{01} = 3.439$, i.e., moderate evidence for the H₀).

540

FIGURE 4

541 3.1 tACS_{triangle} over V1 modulates visual contrast threshold

542 In the first experiment, we investigated whether tACS_{triangle} modulates the visual contrast 543 detection when applied to V1. We measured VCT during tACS_{triangle} at intensities of 0.75, 1, 544 to 1.5mA peak-to-baseline versus no tACS control condition. We found a general decrease in VCT (F_{(3, 81)} = 3.41, p = 0.021, η_p^2 = 0.11, BF_{01} = 0.498) reflecting improved contrast 545 546 sensitivity during tACS_{triangle} (Figure 5A). Post hoc comparisons revealed that 0.75mA and 547 1mA stimulation were most effective in boosting contrast processing at a group level, which 548 differed significantly from the no tACS control condition (p = 0.033, mean difference, MD = -549 $6.3 \pm 11.62\%$ and p = 0.024, MD = -6.33 \pm 10.45%, respectively). Neither the main effect of 550 *block* (F_(1, 27) = 2.43, p = 0.13, BF₀₁ = 1.429) nor *tACS*_{triangle}**block* interaction (F_(3, 81) = 1.6, p = 551 0.195) reached significance.

552 When comparing tACS_{triangle}-induced effects between the 1st and 2nd block we found that the 553 maximal behavioral improvement (i.e., maximal tACS_{triangle}-induced lowering of the VCT 554 relative to the no tACS condition) were not significantly different between the 1st (MD = -555 14.64 \pm 12.6%, VCT decrease in 25 out of 28 individuals) and the 2nd block (MD = -15.75 \pm 556 15.73%, VCT decrease in 24 out of 28 individuals; t₍₂₇₎ = 0.604, p = 0.551, BF₀₁ = 4.219), 557 additionally showing that no time effects arose from the first to the second block of 558 measurement.

Next, we defined the optimal ind-tACS_{triangle} for each participant and examined whether its effects can be reproduced. We observed that the ind-tACS_{triangle} determined in 1st block (**Figure 5B**) caused decrease in VCT compared to the no tACS condition when retested within the same experimental session ($t_{(27)} = 1.84$, p = 0.039, BF₀₁ = 0.463, VCT decrease in 18 out of 28 individuals, MD = -5.26 ± 18.23%, **Figure 5C**). Note, that the above analysis does not contain an element of intrinsic circularity because the ind-tACS_{triangle} and the VCT measure were based on independent data sets.

566 The cutaneous sensation control experiment revealed that some of our participants could 567 detect tACS_{triangle} conditions (HR at 0.75mA = $12.5 \pm 25\%$, 1mA = $18.75 \pm 27.74\%$, 1.5mA = 568 41.07 ± 43.68%, mean HR = 24.11 ± 27.34%). We reanalyzed our main outcome parameter 569 by adding sensation detection HRs for each current level as covariates (HRs were z-scored 570 because of non-normal distribution). The main effect of $tACS_{triangle}$ remained significant (F_{(3, 571 τ_{22}) = 3.36, p = 0.023, η_p^2 = 0.12). Moreover, the mean HR of cutaneous sensation detection 572 did not correlate with the average tACS_{triangle}-induced VCT modulation (r = 0.181, p = 0.357, 573 BF₀₁ = 2.835), making it unlikely that transcutaneous sensation was the main driver of our 574 results.}

575

FIGURE 5

576 3.2 tACS_{sine} over V1 modulates visual contrast threshold

577 In the second experiment, we explored the effects of tACS_{sine} applied over V1 on visual 578 contrast detection. VCT was measured during tACS_{sine} at intensities of 0.75, 1, to 1.5mA 579 peak-to-baseline versus no tACS control condition. We observed a general decrease in VCT with increasing tACS_{sine} intensity ($F_{(3, 81)}$ = 4.78, p = 0.004, η_p^2 = 0.15, BF₀₁ = 0.111) reflecting 580 581 improved contrast sensitivity during tACS_{sine}. Post hoc comparisons revealed that the 1mA 582 and 1.5mA stimulation were most effective in enhancing contrast processing, which differed 583 significantly from the no tACS control condition (p = 0.042, MD = -8.04 \pm 13.82% and p = 584 0.008, MD = -6.52 \pm 12.66%, respectively, Figure 6A). There was no main effect of block $(F_{(1, 27)} = 0.02, p = 0.878, BF_{01} = 3.619)$ or *tACS_{sine}*block* interaction $(F_{(3, 81)} = 0.5, p = 0.684)$. 585

586 When comparing tACS_{sine}-induced effects between the 1st and 2nd block we found that the 587 maximal behavioral improvement, defined as maximal tACS_{sine} induced lowering of the VCT 588 were not different between the 1st (MD = -17.78 ± 15.82%, VCT decrease in 25 out of 28 589 individuals) and the 2nd block (MD = -18.37 ± 16.67%, VCT decrease in 22 out of 28 590 individuals; t₍₂₇₎ = 0.95, p = 0.353, BF₀₁ = 3.320). We determined the optimal ind-tACS_{sine} and tested whether its effects can be reproduced. Similar to ind-tACS_{triangle} in experiment 1, the optimal ind-tACS_{sine} determined in 1st block (**Figure 6B**) significantly lowered the VCT compared to the no tACS condition when retested on the independent VCT data set of the 2nd block ($t_{(27)} = 2.59$, p = 0.008, BF₀₁ = 0.157, VCT decrease in 18 out of 28 individuals, MD = -7.85 ± 21.84%, **Figure 6C**).

596 Similarly to experiment 1, we assessed the HR of cutaneous sensation detection (HR at 597 0.75mA = 16.07 ± 27.4%, 1mA = 21.43 ± 30.21%, 1.5mA = 50.89 ± 43.29%, mean HR = 598 29.46 ± 27.36%). We reanalyzed our main outcome parameter by adding mean cutaneous 599 sensation detection HRs as a covariate (HRs were z-scored because of non-normal 600 distribution). The main effect of tACS_{sine} remained significant ($F_{(3, 72)}$ = 4.67, p = 0.005, η_p^2 = 601 0.16). The mean HR of cutaneous sensation did not correlate with the average tACS_{sine}-602 induced VCT modulation (r = -0.12, p = 0.542, BF₀₁ = 3.569). In this experiment we 603 additionally tested phosphenes detection (HRphos at 0.75mA = 3.57 ± 8.91%, 1mA = 5.36 ± 604 12.47%, 1.5mA = 6.25 ± 16.14%, mean HR = 5.06 ± 10.48%). After adding HR_{phos} as 605 covariate (z-scored HR_{phos}), the main effect of $tACS_{sine}$ remained significant (F_(3, 72) = 4.82, p = 0.004, η_p^2 = 0.17). Accordingly, the mean HR of phosphene detection did not correlate with 606 607 the average tACS_{sine}-induced VCT modulation (r = -0.14, p = 0.493, BF₀₁ = 3.405).

608

FIGURE 6

609 **3.3 Comparison of tACS**_{triangle}, tACS_{sine}, and hf-tRNS-induced modulation

First, we tested whether baseline VCT measured during the no tES conditions differed between the experiments (i.e., average baseline VCT in tACS_{triangle}, tACS_{sine}, and hf-tRNS experiments, see **Figure 4**). Bayesian independent samples t-test revealed that the baseline VCT measured in the no tES condition did not differ between experiments ($BF_{01} = 4.869$, i.e., moderate evidence for the H₀). Next, we compared tES-induced modulation effects between experiments (tACS_{triangle}, tACS_{sine} and hf-tRNS experiments, see **Figure 7A**). A Bayesian ANOVA revealed that the general tES-induced modulation did not differ between experiments ($BF_{01} = 8.956$, i.e., moderate evidence for the H₀), suggesting that all three stimulation types were equally effective in lowering VCT.

620 Finally, we assessed the strength of the tES-induced effects on VCT across tACS_{triangle}, 621 tACS_{sine} and hf-tRNS experiments defined as paired Cohen's d bootstrapped sampling 622 distributions (see Figure 7B). We found comparable (small) effects of significant differences 623 between no tES baseline VCT and averaged VCT in active tES conditions in all experiments 624 using the two-sided permutation t-test [in tACS_{triangle} d = -0.17 (95.0%CI -0.284; -0.0698) p = 625 0.0034; in tACS_{sine} d = -0.242 (95.0%CI -0.444; -0.103), p = 0.0016; in hf-tRNS d = -0.249 626 (95.0%Cl -0.433; -0.088) p = 0.0092]. The effect sizes and Cls are reported above as: effect 627 size (CI width lower bound; upper bound).

628

FIGURE 7

629 4. Discussion

630 Theoretical modelling shows that adding a deterministic, high-frequency sinusoidal signal 631 instead of stochastic noise could lead to signal enhancement due to resonance, according to 632 the DAR mechanism. Our experimental proof-of-concept study revealed, that stimulation of 633 V1 with a deterministic tACS signal instead of stochastic noise leads to signal enhancement 634 in visual processing. We measured visual contrast sensitivity during tACS_{triangle} and tACS_{sine}. 635 On the group level, we found consistent tACS_{triangle}- and tACS_{sine}-induced decrease in VCT, 636 reflecting enhancement in visual contrast processing during V1 stimulation (Figure 5A, 637 Figure 6A). The online modulation effects of individually optimized tACS_{triangle} and tACS_{sine} 638 intensities (Figure 5B, Figure 6B) were replicated on the independent VCT data (Figure

5C, Figure 6C). Finally, we demonstrated that the effects of deterministic stimulation on VCT
are comparable to stochastic stimulation of V1 with hf-tRNS (Figure 7AB).

641 **4.1 tACS with triangle and sine waveform improve visual sensitivity**

642 Our findings provide the first proof of concept that the deterministic $tACS_{triangle}$ and $tACS_{sine}$ 643 delivered to V1 can modulate visual contrast sensitivity. Across two experiments we showed 644 that the modulatory effects of tACS on visual sensitivity are not waveform specific, as both 645 $tACS_{triangle}$ and $tACS_{sine}$ induced significant decrease in VCT (**Figure 5A**, **Figure 6A**).

One of the main characteristics of SR-like effects is the optimal intensity of noise, which is required in order to yield the improved performance (5, 15). Here, we did not observe an excessive level of tACS that would be detrimental for visual processing (**Figure 5A**, **Figure 64**9 **6A**). This is consistent with our predictions that, in line with DAR (see mathematical predictions in 1.2.1), adding high frequency deterministic signal should result in a noise-free output where the detection processing is not disturbed by random stimulation effects.

Similar to other studies investigating resonance-like effects (28, 39), our results have revealed large variability among participants in terms of the optimal intensity resulting in the strongest modulation of visual contrast sensitivity (**Figure 5B**, **Figure 6B**). However, consistent with the effects of tRNS-induced online modulation of contrast processing in V1 shown previously (28, 39), the effects of individualized tACS intensity were replicated on the independent VCT data set collected within the same experimental session (**Figure 5C**, **Figure 6C**), suggesting consistent beneficial influence of tACS on signal enhancement.

We implemented several control measures to test whether the improvement in visual processing was driven by effective stimulation of V1 rather than any unspecific effects of tACS. We applied an anesthetic cream to numb potential stimulation-induced cutaneous sensation on the scalp (50, 51). While the anesthetic cream numbs the skin and reduces the cutaneous sensations resulting from tACS, it does not eliminate them completely in all

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664 individuals. The control cutaneous sensation detection assessment in the current study 665 showed that some participants could accurately detect tACS, and that the mean detection 666 rate was rather low (mean HR = 24.11 \pm 27.34% in tACS_{triangle} and mean HR = 29.46 \pm 667 27.36% in tACS_{sine}). Cutaneous sensation and phosphenes detection (also very low, mean 668 $HR_{phos} = 5.06 \pm 10.48\%$) did not correlate with the average tACS-induced VCT modulation 669 neither in tACS_{triangle}, nor tACS_{sine} experiment. Moreover, stimulation effects remained 670 significant in the additional analysis using tactile or phosphene sensation detection during 671 tACS_{triangle} and tACS_{sine} as covariate.

672 While tACS_{sine} is a well-established and frequently used non-invasive brain stimulation 673 method, high frequency tACS_{sine} is less common. The effects of 80Hz tACS_{sine} were 674 sporadically tested in the past using physiological and behavioral paradigms. Ten minutes of 675 140Hz tACS_{sine} was shown to increase primary motor cortex (M1) excitability as measured 676 by transcranial magnetic stimulation-elicited motor evoked potentials during and for up to 1h 677 after stimulation. Control experiments with sham and 80Hz stimulation did not show any 678 effect, and 250Hz stimulation was less efficient, with a delayed excitability induction and 679 reduced duration (60). The researchers postulated that the changes in corticospinal 680 excitability result from externally applied high frequency oscillation in the ripple range (140Hz 681 corresponding to middle, 80Hz lower and 250Hz upper border) that interfere with ongoing 682 oscillations and neuronal activity in the brain (60). We can, however, not directly translate 683 the effects of tACS_{sine} of M1 to our stimulation of V1. Additionally, the stimulation effects 684 observed in our study are likely reflecting acute modulation of contrast processing, as 685 stimulation was only applied for short intervals (2 s) always interleaved with control (no 686 tACS) condition. Thus, it is possible that even though 80Hz stimulation did not lead to long 687 term effects in cortical excitability it can still affect cortical processes acutely.

688 In the visual domain, 1.5mA high-frequency tACS_{sine} was applied to V1 for 15-45min in a 689 study investigating the effect of covert spatial attention on contrast sensitivity and contrast 690 discrimination (61). That study found that contrast discrimination thresholds decreased 691 significantly during 60Hz tACS_{sine}, but not during 40 and 80Hz stimulation. This previous 692 study used, however, different visual stimuli than that utilized here, i.e., a random dot 693 pattern. Moreover, they used a more complicated behavioral paradigm, where contrast-694 discrimination thresholds were tested using two attention conditions, i.e., with or without a 695 peripheral cue, as the study goal was to explore the influence of attentional processes on 696 visual tasks. One tACS mechanism that has been tested in the visual domain is the 697 reduction of adaptation (62, 63). More specifically, a seminal study has shown that 10Hz 698 tACS reduces sensory adaptation in a visual motion perception task (62). Since sensory 699 adaptation increases thresholds for detection, potentially reducing adaptation during tACS 700 could result in decreased thresholds. However, the aforementioned study design differed 701 substantially from the one presented here. Kar and Krekelberg (62) used a 40s adaptor 702 stimulus to induce adaptation while our stimuli were presented for only 40ms (i.e., 3 703 magnitudes shorter). This is relevant because it was previously shown that adaptation gets 704 stronger and lasts longer with increasing adaptation duration (64). Moreover, Kar and 705 Krekelberg (62) used 10Hz tACS which was substantially lower than the one exploited in our 706 experiments (i.e., 80Hz), making direct comparisons difficult. Overall, additional experiments 707 would be required to test whether our results could be explained by a reduction of visual 708 adaptation.

709 Even though the vast majority of tACS studies to date have used a sinusoidal waveform, an 710 alternating current does not have to be sinusoidal, since it can take any arbitrary waveform 711 such as rectangular wave (65), pulsed (66), or sawtooth (67). Dowsett and Herrmann (2016) 712 investigated the effects of sinusoidal and sawtooth wave tACS on individual endogenous 713 alpha-power enhancement. They observed alpha oscillations enhancement both during and 714 after sawtooth stimulation. The effect seemed to depend on the shape of the sawtooth, as 715 they found that positive, but not negative, ramp sawtooth significantly enhanced alpha power 716 during stimulation relative to sham. They postulated that a sudden, instantaneous change in

current might be more effective than a sinusoidal current in increasing the probability of neurons firing. In this regard, Fröhlich and McCormick (Supplementary Material in (68)) demonstrated that ramps of increasing voltage with a steeper gradient resulted in increased neural firing in vitro, relative to ramps with a low gradient but reaching the same maximum voltage. This suggests that it is not only the total amount of current but also the rate of change of current can modulate neural firing. Note, that triangle waveform has a faster rate of change of current than the sine wave.

724 Although we postulate that the effect of tACS on VCT in our study results from resonance-725 like mechanism, this is not the only potential mechanism. Importantly, the commonly 726 accepted mechanism of action of tACS is that it entrains action potential firing, and thus 727 neural oscillations (69). Entrainment effect anticipates a monotonic relationship between the 728 tACS effect and intensity, where increasing stimulation intensity results in greater effects for 729 stimulation waveforms that are tuned to the endogenous oscillation (70). The effects of tACS 730 in regard to induced brain oscillations seem to depend on the stimulation duration (71). 731 Although the entrainment after-effects were observed after tACS had been delivered for 732 several minutes (71), short stimulation of 1s did not produce after-effects on amplitude or 733 phase of the electroencephalogram (72). Moreover, in the study investigating the effects of 734 intermittent alpha tACS of either 3 or 8 s, the after-effects were found only for the 8-s 735 condition (73). The authors excluded entrainment as potential underlying mechanism and 736 postulated plasticity-related changes as the responsible mechanism for the observed after-737 effects. Here, we used very brief stimulation of 2s tACS per trial, a duration seemingly too 738 short to induce the entrainment effects on cortical processing.

Furthermore, it was postulated that a very small amount of applied electric field can bias
spike timing or spike probability when a neuron nears the threshold of spike generation (74).
Accordingly, it was shown that although entrainment effects can arise at field strengths <0.5
mV/mm, physiological effects are more pronounced for higher intensities (around 1mV/mm),

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743 according to intracranial recordings in awake nonhuman primates (75). These values are 744 well above the simulated induced electric field in our study (around 0.2 mV/mm, see Figure 745 3). Further studies are required to fully disentangle the underlying neuronal effects of tACS 746 driving the enhancement in visual detection. To exclude the influence of entrainment on VCT 747 modulation a jittered tACS protocol could be employed. A paradigm using stimulation of 748 jittered flickering light, where instead of a rhythmic flicker, inter stimulus intervals of the 749 square wave were jittered with a maximum of ± 60%, was shown to fail in inducing rhythmic 750 brain response (76). If a jittered tACS of V1 would still influence contrast sensitivity we could 751 assume the non-entrainment origin of the effect.

752 **4.2 Comparison of tACS**_{triangle}, tACS_{sine}, and hf-tRNS

753 In the phenomenon of SR, random noise added to a non-linear system can increase its 754 responsiveness towards weak subthreshold stimuli. One aim in the present study was to 755 explore whether a deterministic and periodic signal can substitute stochastic noise and still 756 lead to response enhancement in a threshold-based stochastic resonator. The DAR 757 characteristics of high-frequency deterministic signal might offer a noise-free output, thus 758 additionally increasing SNR. We proposed the following testable hypotheses (i) tACS_{triangle} 759 will have a larger resonance-like effect compared to hf-tRNS, (ii) tACS_{sine} will have less effect 760 than tACS_{triangle}, due to the loss of waveform linearity. We found enhancement effects of both 761 tACS_{triangle} vs tACS_{sine} (Figure 5A, Figure 6A), however to test whether these effects are 762 indeed superior to stochastic stimulation, we directly compared the VCT modulation induced 763 by tACS_{triangle}, tACS_{sine} and hf-tRNS (**Figure 7**). The baseline contrast sensitivity between the 764 compared experiments was not different (Figure 4). Counter to our hypothesis, the noise-765 free tACS did not result in stronger contrast sensitivity enhancement, as average VCT 766 modulation did not differ between the three stimulation conditions, as confirmed by Bayesian 767 analysis (Figure 7A). Accordingly, the effects sizes of all three stimulation types were comparable (Figure 7B). Therefore, we showed that both deterministic and stochastic high frequency stimulations were equally effective in inducing resonance-like effects.

770 In real life (in comparison to mathematical simulations) neural processing is intrinsically 771 noisy. How this intrinsic noise interacts with the applied noise/SR signal will have 772 implications for the validity of our mathematical model. The task we used is a 4AFC 773 discrimination task, which means that when tRNS is added to the neurons that there must be 774 a distinction between 3 noisy locations and 1 signal and noise location. How added noise 775 influences this comparison remains unresolved and an area for further investigation. To 776 investigate this, a two-sided tRNS experiment could be run where noise is added to the left 777 and/or right V1 (or S1) to explore the influence of more noise added to the system where the 778 signal is not present versus when it is present. This might give some information on how the 779 brain interacts with signal, added noise and intrinsic noise compared to intrinsic noise only 780 on a discrimination task.

781 4.3 Conclusions

The present study provides the first evidence for resonance-like neural signal enhancement without adding a stochastic noise component. We showed that 'deterministic' 80Hz-tACS and 'stochastic' hf-tRNS are equally effective in enhancing visual contrast detection. In the range of commonly used intensities of tES to induce SR, tACS did not result in detrimental effects related to excessive interference signal, thus providing increased SNR in all tested intensities, according to DAR predictions. These findings shed a new light on the effects induced by both 80 Hz tACS and hf-tRNS, and their underlying mechanisms.

The optical excitations in this work were square wave signals (i.e., the visual stimulus was switched on for 40 ms). Open theoretical and related experimental questions are: What is the role of the finite time it takes for the square wave signal to reach its amplitude? What if

- the signal has a different periodicity? Possible interrelations between the duration of the
- 793 pulses and the inter-pulse intervals?

GLOSSARY

800

795 Stochastic Resonance (SR) – certain nonlinear systems show improved signal transfer in

the presence of high-frequency additive noise. It is an amplitude resonance because there is

an optimal noise (root mean square) amplitude for the best transfer.

- 798 **Threshold Elements (TE)** a device with threshold-based nonlinearity.
- 799 Level Crossing Detector (LCD) a threshold element that produces a short uniform spike

signal at its output whenever the input signal crosses the threshold level. There are

- variations depending on what type of crossing (up, down, or both) triggers a spike.
- 802 **Comparator** a threshold element that produces zero output value whenever the input
- signal is below the threshold and a non-zero U_H value otherwise.
- 804 Signal strength (SS) the mean-square amplitude of the signal.

805 **Signal-to-noise-ratio (SNR)** - the ratio of the mean-square amplitudes of signal and noise.

- 806 **Information entropy** the maximum of the useful information that an unknown message
- 807 with a given size can contain.
- 808 **Shannon information channel capacity** the maximum bit rate that a (typically noisy) 809 information channel can effectively transfer.
- 810 **Deterministic Amplitude Resonance (DAR)** a device that, similarly to SR, show improved 811 signal transfer in the presence of high-frequency, additive, deterministic, carrier-wave. It is 812 an amplitude resonance because there is optimal carrier-wave amplitude for the best 813 transfer.
- 814 **Periodic carrier-waves** carrier waves that are periodic time functions.
- 815 **Triangle wave** periodic carrier-wave with straight lines of the rising and the falling sections.
- 816 Sine wave sinusoidal carrier-wave

817 **Transcranial Electric Stimulation (tES)** – noninvasive brain stimulation technique, which 818 applies weak, painless electrical currents to the scalp (current intensities of \sim 1–2 mA), to 819 modulate brain function (77–80).

820 Transcranial Alternating Current Stimulation (tACS) - a subtype of tES characterized by
821 biphasic, alternating electric currents applied (69, 81).

Transcranial Random Noise Stimulation (tRNS) - a subtype of tES whereby currents are
 randomly drawn from a predefined range of intensities and frequencies (26, 82, 83).

Visual contrast detection threshold (VCT) – criterion reflecting the level of task performance accuracy. Here, the detection threshold corresponds to the contrast intensity of presented visual stimuli that was accurately detected with 50% accuracy (see Figure 3).

QUEST staircase procedure - a method used in psychophysical research to estimate threshold of a psychometric function (47). In this maximum-likelihood adaptive procedure, information from all trials in an experiment are considered to determine a threshold (47, 48). Here, QUEST method was used to estimate the visual contrast detection threshold for each participant.

832 Four-alternative forced choice (4-AFC) visual task - design of a discrimination task in 833 psychophysical experiments, where participant is forced to choose one out of four possible 834 responses. In contrast to methods requiring a 'yes/no' response, forced-choice methods 835 characterize with higher accuracy of the measured psychophysical property (45). Here, the 836 weak visual stimulus was presented with different intensities in one of 4 quadrants on the 837 screen and participants were asked to select in which one it appeared in each trial. Based on 838 the accuracy of those responses we estimated their contrast detection threshold using 839 QUEST procedure.

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1094

1095 Figures Captions

1096 Figure 1 Deterministic transfer of sub-threshold binary signal through simple threshold-based stochastic 1097 resonators with a Threshold Element (TE: either a Level Crossing Detector (LCD) or a Comparator) and an 1098 additive triangle wave at the input. Note: the classical threshold-based stochastic resonators contain the same 1099 hardware elements except the triangle wave that is substituted by a Gaussian random noise. The role of the Low-1100 pass Filter is to reduce the amount of irrelevant high-frequency products created by the carrier wave. If those 1101 irrelevant high-frequency products are not disturbing, the Low-pass Filter can be omitted. Upper part: the sub-1102 threshold binary signal is unable to trigger the TE thus the output signal is steadily zero. Lower part: an additive, 1103 triangle wave (carrier-wave) assists the signal to reach the threshold thus it carries the binary signal over the TE. 1104 The Low-pass filter takes a short time average in order to smooth out the high-frequency components. For high-1105 fidelity transfer, to avoid problems caused by delays or phase shifts, the frequency of the carrier-wave must be 1106 much greater than that of the binary signal. In the old stochastic resonance schemes, the carrier-wave was a 1107 noise that caused a non-deterministic component (noise) and finite SNR at the output. The new system is purely 1108 deterministic, and its SNR is infinite. Moreover, if the signal is "analog" (continuum amplitude values), the triangle 1109 wave with comparator as TE guarantees a linear transfer of the signal provided the threshold level is between the 1110 minimum and the maximum of the sum of the signal and the carrier-wave, see in (ii) below. $U_{\rm th}$ = threshold, $U_{\rm s}$ =

1111 signal, U_t = noise, U_{lcd} = LCD output signal, U_c = comparator output signal, U_{LPF} = signal after low-pass filtering.

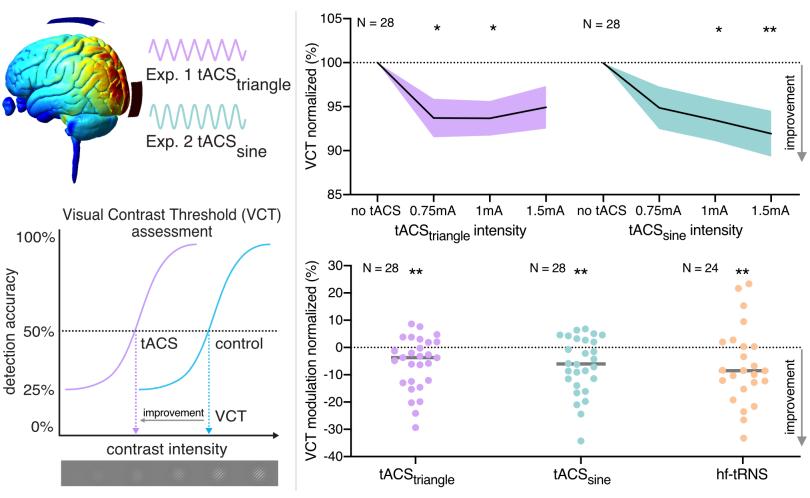
1112 Figure 2 The triangle wave vs. the threshold (U_{th}).

1113 Figure 3 Experimental design. A. Example trial of 4-alternative forced choice task measuring visual contrast 1114 detection threshold (VCT). tACS was delivered for 2 s around the Gabor patch presentation. B. tACS electrodes 1115 montage targeting V1 and simulation of the induced electric field in the brain. C. Example of dose-response 1116 psychometric curves and the VCT for the 50% detection accuracy level. We hypothesize that the VCT will be 1117 lower (indicating better contrast detection performance of the participant) in one of the tACS conditions (violet) 1118 than in the no tACS control condition (blue). D. The order of measurements within each experiment. Each 1119 experimental session consisted of application of an anesthetic cream, followed by task training, familiarization 1120 protocol, and two independent VCT assessments in 4 interleaved tACS conditions (as specified in A).

Figure 4 Average baseline VCT measured in the no tES conditions in tACS_{triangle} N=28 (16 females, 12 males),
 tACS_{sine} N=28 (20 females, 8 males), hf-tRNS experiments N=24 (16 females, 8 males). VCT was assessed for
 stimuli presented with contrast intensity ranging from 0 to 1. Blue lines indicate mean, gray dots indicate single
 subject data. BF₀₁ = 4.869, i.e., moderate evidence for the H₀.

Figure 5 The effect of tACS_{triangle} on VCT measured in experiment 1. VCT was assessed for stimuli presented with contrast intensity ranging from 0 to 1. **A.** Effect of tACS_{triangle} on VCT on a group level measured across 1st and 2nd blocks. Decrease in VCT reflects improvement of visual contrast sensitivity. VCT in tACS_{triangle} conditions normalized to the no stimulation baseline. All data mean \pm SE; *p < 0.05, rmANOVA **B.** Individually defined optimal tACS_{triangle} based on behavioral performance during the 1st block. **C.** Detection improvement effects of individualized tACS_{triangle} (selected based on block 1) measured on the independent VCT data of block 2. Gray dots indicate single subject data; *p < 0.05, t-test for dependent measures. N=28 (16 females, 12 males).

Figure 6 The effect of tACS_{sine} on VCT measured in experiment 2. VCT was assessed for stimuli presented with contrast intensity ranging from 0 to 1. **A**. Effect of tACS_{sine} on VCT on a group level measured across 1st and 2nd blocks. Decrease in VCT reflects improvement of visual contrast sensitivity. VCT in tACS_{sine} conditions normalized to the no stimulation baseline. All data mean \pm SE; *p < 0.05, **p < 0.01, rmANOVA. **B**. Individually defined optimal tACS_{sine} based on behavioral performance during the 1st block. **C**. Detection improvement effects of individualized tACS_{sine} (selected based on block 1) measured on the independent VCT data of block 2. Gray dots indicate single subject data; **p < 0.01, t-test for dependent measures. N=28 (20 females, 8 males). 1139 Figure 7 Comparison of tACS_{triangle} (N=28; 16 females, 12 males), tACS_{sine} (N=28; 20 females, 8 males), and hf-1140 tRNS-induced modulation (N=24; 16 females, 8 males). A. VCT modulation induced by tACS_{triangle}, tACS_{sine}, hf-1141 tRNS. The general modulation of VCT induced by tES was calculated as mean of all active tES conditions from 1st and 2nd blocks normalized to baseline no tES condition in each experiment. Decrease in VCT reflects 1142 1143 improvement of visual contrast sensitivity. **p < 0.01, two-sided permutation t-test. B. The paired Cohen's d for 3 1144 comparisons shown in the Cumming estimation plot. Each paired mean difference is plotted as a bootstrap 1145 sampling distribution. Mean differences are depicted as dots, 95% confidence intervals are indicated by the ends 1146 of the vertical error bars.



The 'deterministic' tACS (with triangle- and sine-waveforms) and 'stochastic' hf-tRNS are equally effective in enhancing visual contrast detection

