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Research Articles: Behavioral/Cognitive

## The assimilation of novel information into schemata and its efficient consolidation

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1	The assimilation of novel information into schemata and its
2	efficient consolidation
3	Short title: Assimilation and consolidation of schema-related facts
4	Tobias Sommer <sup>1†</sup> , Nora Hennies <sup>1</sup> , Penelope A. Lewis <sup>2*</sup> , Arjen Alink <sup>1*</sup>
5	
6	<sup>1</sup> Institute of Systems Neuroscience, University Medical Center Hamburg-Eppendorf, 20246 Hamburg,
7	Germany
8	<sup>2</sup> CUBRIC, School of Psychology, Cardiff University, Cardiff CF24 4HQ, UK
9	<sup>†</sup> corresponding author: <u>tsommer@uke.de</u>
10	* these author contributed equally
11	
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### 19 Abstract

20 Schemata enhance memory formation for related novel information. This is true even 21 when this information is neutral with respect to schema-driven expectations. This 22 assimilation of novel information into schemata has been attributed to more effective 23 organizational processing that leads to more referential connections with the activated 24 associative schema network. Animal data suggest that systems consolidation of novel 25 assimilated information is also accelerated. In the current study, we used both multivariate 26 and univariate fMRI analyses to provide further support for these proposals and to 27 elucidate the neural underpinning of these processes. 28 Participants (5 male) over-28 learned fictitious schemata for seven weeks and then encoded novel related and control 29 facts in the scanner. These facts were retrieved both immediately and two weeks later, 30 also in the scanner. Our results conceptually replicate previous findings with respect to 31 enhanced vmPFC-hippocampus coupling during encoding of novel related information and 32 point to a prior knowledge-effect that is distinct from situations where novel information is 33 experienced as congruent or incongruent with a schema. Moreover, the combination of 34 both multi- and univariate results further specified the proposed contributions of the 35 vmPFC, precuneus and angular gyrus-network to the more efficient encoding of schema-36 related information. In addition, our data provide further evidence for more efficient 37 systems consolidation of such novel schema-related and potentially assimilated 38 information.

### 39

### 40 Significance Statement

41 Our prior-knowledge in a certain domain, often termed schema, heavily influences whether 42 and how we form memories for novel information that can be related to them. The results 43 of the current study show how a ventromedial prefrontal-precuneal-angular network 44 contributes to the more efficient encoding of novel related information. Furthermore, the 45 observed increase in prefrontal-hippocampal coupling during this process points to a 46 critical distinction from the previously described mechanisms supporting the encoding of 47 information that is experienced as congruent with schema-driven expectations. In addition, 48 we find further support for the proposal based on animal data that prior-knowledge

49 enhances also the consolidation of schema-related information.

### 50 Introduction

51 Whether and how we encode, consolidate and later retrieve novel information is heavily 52 influenced by our prior-knowledge (Alba and Hasher, 1983; Gilboa and Marlatte, 2017). 53 The impact of prior-knowledge on memory has been studied in the framework of schemata 54 (Bartlett, 1932) which are previously acquired and continuously developing associative 55 networks (Ghosh and Gilboa, 2014).

56 A prominent line of research on the prior-knowledge effect investigates how the 57 congruency of information with schema-driven expectations affects memory, e.g. 58 remembering a palm-tree vs. a polar bear at a beach (van Kesteren et al., 2012; Greve et 59 al., 2019). However, prior-knowledge enhances also memory for novel, previously 60 unknown - and hence expectation-neutral - information that can be related to it, for 61 instance when we learn new facts in our academic field (Brandt et al., 2005; Witherby and 62 Carpenter, 2021). The vmPFC together with the ventral precuneus/retrosplenial cortex 63 (vPC/RSC) are involved also in this effect of prior-knowledge where the contribution of the 64 hippocampus and its coupling with the vmPFC remain so far inconsistent across studies 65 (Tse et al., 2011; van Kesteren et al., 2014; Brod et al., 2016; Liu et al., 2016; Sommer, 66 2017).

The mnestic advantage for novel schema-related but expectation-neutral information has been attributed to the activation of schema-knowledge. This allows more effective organizational processing leading to assimilation into the associative structure (Ericsson and Kintsch, 1995). In particular, the newly encoded information might be integrated via spreading activations into the existing associative network resulting in referential connections and the association with appropriate retrieval cues (Long and Prat, 2002).

74 These processes underlying assimilation would not necessarily result in enhanced 75 mean activity in the involved brain areas compared to encoding of schema-unrelated 76 information. The first goal of the current study was therefore to find support for such 77 processes by using multivariate representational similarity analyses (RSA). In particular, 78 after participants had acquired knowledge of experimental schemata (Fig. 1 A) we 79 contrasted the similarity of activity patterns during three encoding rounds of novel schema-80 related with those of tightly matched control facts. By this means we tested the hypothesis 81 that the rapid integration of schema-related facts into the associative structures results in more consistent representation across encoding rounds (i.e. greater pattern robustness; 82 83 Xue et al., 2010; Bruett et al., 2020). Moreover, we tested whether the more effective

organizational processing would be reflected in more consistent encoding operations
across novel schema-related facts. Finally, we tested the hypothesis that assimilation
should be evident in higher similarity between encoding of novel schema-related facts and
the retrieval of overlearned schema-knowledge.

88 A second goal of the current study was to provide further evidence with respect to 89 hippocampal-vmPFC coupling during encoding of novel schema-related but expectation-90 neutral information. In the two studies observing stronger coupling participants encoded 91 arbitrary associations (Liu et al., 2016; Sommer, 2017) whereas in the third students 92 learned real facts, i.e. associations of an already taught with a new term, in their or another 93 discipline (van Kesteren et al., 2014). The diverging results might be caused by the fact 94 that in the other discipline both terms forming a new fact were novel (Carpenter et al., 95 2018) or that only in the latter study meaningful information was encoded. Both factors 96 were addressed in the current paradigm with the goal to better understand hippocampal-97 vmPFC coupling during encoding of novel schema-related information.

98 A series of animal studies showed that prior-knowledge not only influences 99 encoding but also accelerates systems consolidation of novel schema-related expectation-100 neutral information, i.e. randomly paired flavor-location associations (Tse et al., 2007). In 101 the aforementioned study using a similar design, we showed that also in humans schema-102 related information might be more rapidly transferred from hippocampal to cortical retrieval 103 (Sommer, 2017). Using meaningful knowledge structures we also showed that nightly 104 replay might underlie this effect (Hennies et al., 2016). A third goal of the current study was 105 therefore to use those meaningful associative structures to conceptually replicate our 106 previous findings and to provide further evidence that novel schema-related expectation-107 neutral information is transferred more rapidly to neocortical retrieval. To test this 108 hypothesis, participants' memory for novel facts was tested in the MRI scanner 109 immediately after encoding and 2 weeks later. 110 Methods

### 111 Participants

112 Thirty-two native Germans (mean age: 25.90 years; SD: 3.71 years; 6 males; randomly

- 113 assigned to one of the two schemata) participated in the study. All had normal or
- 114 corrected-to-normal vision, and no history of neurological and psychiatric disorders.
- 115 Participants were required to have no knowledge beyond that of basic schooling at the
- 116 primary level in the two schema categories (arthropods and cell biology) and no particular
- 117 interest in biology, medicine, chemistry, and zoology. Informed consent was obtained from

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- 118 all participants before the study, which was approved by the Ethics Committee of the
- 119 Hamburg Medical Association. The first participant could not be analyzed due to data loss.
- 120 Two participants dropped out of the study during the schema-knowledge acquisition. For
- 121 one participant, retrieval results are missing due to technical failure, leaving 28 participants
- 122 (5 male, 14 for each of the two schemata).

### 123 Stimuli

- 124 The aim was to experimentally construct two fictitious schemata that fulfilled the four
- 125 previously identified criteria, i.e. an associative network structure, basis on multiple
- 126 episodes, lack of unit detail, and adaptability, (Ghosh and Gilboa, 2014) and the
- 127 neuroscientific definition, i.e. consolidated cortical representation (van Kesteren et al.,
- 128 2012). At the same time, we aimed to minimize previously described confounds when pre-
- 129 existing, e.g. academic or other expert, knowledge is used: i) higher
- 130 curiosity/interest/motivation to learn novel facts in the domain of expertise and ii) the
- 131 possibility that novel schema-related information had been known before (Witherby and
- 132 Carpenter, 2021). Moreover, in the area of expertise participants have a higher familiarity
- 133 with the names of the concepts the novel information has to be associated with, e.g. when
- 134 "blossom end rot" has to be associated with calcium deficiency the term "blossom end rot"
- 135 is for experts familiar but novel for control participants (Carpenter et al., 2018). Such
- 136 greater cue familiarity results in a processing advantage and in differences in (meta-)
- 137 memory (Chua et al., 2012). Importantly, as the goal of the current study was to
- 138 characterize the effects of prior-knowledge on memory due to more effective
- 139 organizational processing and not to in-/congruency, we aimed to minimize the influence of
- schema driven expectations with respect to the novel facts. However, we also aimed to
- 141 use not arbitrary associative structures as in our previous and the animal studies (Tse et
- 142 al., 2007, 2011; Sommer, 2017) but more ecological valid, i.e. meaningful experimental
- 143 schemata. Novel facts related to such meaningful knowledge structures can be only
- 144 neutral with respect to specific expectations based on prior experience (as for instance
- 145 palm-trees and not polar bears are expected at a beach scenery) but are still consistent
- 146 with more general knowledge. To stay with the above example, even if there is no specific
- 147 schema-driven expectation for the cause for "blossom end rot" the novel fact "calcium
- 148 deficiency" is similar to many other possibilities, e.g. sodium deficiency or a fungal attack,
- 149 that are generally consistent with the student's knowledge about phytology. The absence
- 150 of specific schema-driven expectations results at the same time in more detailed, less
- 151 generalized and gist-like memories (Tse et al., 2007, 2007; Sommer, 2017) which diverges

152 from the more typical schema studies on the effect of schema-congruency where higher

153 false alarm rates are observed (Rojahn and Pettigrew, 1992; van Kesteren et al., 2012).

### 154 Schemata

155 The stimuli from our previous study were translated into German and modified for the 156 purpose of the current study (Hennies et al., 2016). The schemata were constructed with 157 four hierarchical levels (Fig.1 A). The two schemata were parallel in structure and 158 contained facts at each level. In particular, each schema comprised facts about the 159 category (arthropods or cells), its two sub-categories (ant and crabs, cell-types and 160 organelles), their three families each (e.g. symbiotic, hunter and weaver ants) and detailed 161 characteristics of the 12 individual exemplars (two in each family). The detailed facts 162 (approximately 25 facts) about anatomy, habitat, food preferences, and behavioral 163 characteristics for each of the 12 exemplars comprised the main part of the 'arthropods'-164 schema. For the 'cells'-schema a matched number of facts for each of the 12 exemplars 165 existed (see Hennies et al., 2016 for family labels and pictures of the exemplars of the 166 'cells'-schema). In addition, each exemplar was presented on 12 different pictures. Novel 167 names were invented for all exemplars in order to avoid large differences in length and 168 complexity of the names. The whole hierarchical associative structure and all facts, i.e. the 169 pictures of the exemplars, related to its nodes and elements were considered as schema -170 similar to studies employing pre-existing academic or trivial real-world knowledge.

### 171 Novel, schema-related facts

172 For each of the exemplars of both schemata six additional facts were created. Each facts 173 existed in two, equally likely alternatives (e.g. Styga is 2 or 4 cms long, NIV contains 174 copper or nickel, Fig. 1B). One version, which was randomly chosen for each participant, 175 was encoded as novel schema-related (SR) fact, the other one served as lure for the 2-176 alterantive forced-choice memory tests. This design ensured that participants could not 177 quess the correct response based on their schema-knowledge and that the novel schema-178 related facts were indifferent with respect to schema congruency and expectations. The 179 SR facts for the 'arthropods'-schema served as non-schema related (NS) control facts for 180 the participants who acquired knowledge for the 'cells'-schema and vice versa. The facts 181 were kept vague to minimize participants guessing about what type of exemplars the 182 control facts were about. The novel facts for both schemata were counterbalanced for the 183 number of words, number of syllables, and numerical values. None of the facts were

longer than eight words or 14 syllables so that all facts could easily be read within thepresentation time.

### 186 Procedure

This experiment was realized with custom-written scripts using Cogent 2000, developed by
the Cogent 2000 team at the Functional Imaging Laboratory and the Institute for Cognitive
Neuroscience (University College, London).

### 190 Acquisition of structured, associative schema-knowledge

191 In a first session in the institute, participants were randomly assigned to one of the two 192 schemata and performed a test on their prior-knowledge in that category. This pretest 193 involved one picture and 6 statements about each of the 12 category exemplars (using 194 their real not the invented names). When a picture was presented the participant could 195 select the name of the exemplar from 6 response options or select a "?" to indicate not 196 knowing. In response to the statements participants could select "true", "false" or "?". All 197 questions were presented without a time limit. No subjects achieved more than the cut-off 198 of 20% correct responses and therefore all could take part in the experiment.

199 The acquisition of schema-knowledge started after the pre-test in the first session 200 and involved seven more sessions in the institute, which were separated by seven days. 201 There were very few exceptions, when participants could not make that day, they came 202 within around 2 days. Sessions were  $\sim$ 1 h long (only session 1 took 2 h), but this could 203 vary as participants completed most tasks in a self-paced way. Between the sessions at 204 the institute, participants deepened their schema-knowledge by working through 205 homework. The first two sessions in the institute and the corresponding homework will be 206 described more in detail in order to give an impression how acquisition of structured 207 schema-knowledge was achieved.

208 In the first session participants started to learn general background information 209 about their schema (on the category and sub-category level), that was presented self-210 paced on the computer. Participant's assigned to the 'arthropods'-schema learned for 211 instance that arthropods have an exoskeleton, a segmented body, compound eyes, that 212 the exoskeletons are based on chitin and vary with respect to their stability, that ants have 213 two antenna, two mandibles, live in colonies etc. Participants did at the end of the first 214 session a multiple-choice test on the acquired knowledge with 32 questions with 87.7% 215 (±10%) accuracy (Fig. 1 B). In the following homework the schema facts from session 1 216 and additional facts were presented as reading material. In addition, the families and

exemplars were introduced including the first facts about both hierarchy levels. Moreover,
each exemplar was presented on 2 pictures from different perspective. Participants were
asked to work through the material and to answer open questions in writing to facilitate
and deepen the learning. They were told, that their answers were evaluated in the next
session in the institute.

In the second session in the institute, participants were asked to free recall the facts 222 223 about the exemplars followed by the multiple choice and ten open questions about the 224 schema background. They then repeated learning the background facts followed again by 225 multiple choice and open questions. In the following homework, they repeated the general 226 background facts as well as learning more facts about the families and exemplars by 227 reading and responding to open question. After this homework they had learned all 228 relevant schema-related facts the first time. From there on, this knowledge was repeated 229 and deepened.

In the remaining 6 sessions in the institute and the 5 homework sessions between them participants continued in the same way to recapitulate the facts about all hierarchy levels of their schema, including 12 pictures of each exemplar and their knowledge was tested using various multiple choice and open questions as well as picture naming tests. At the end of each session participants' schema-knowledge was assessed with multiple choice tests for which they did not receive feedback. Based on their performance in these tests the homework sessions were individually adjusted.

237 Participants showed close to ceiling performance in the multiple-choice and picture 238 naming tests in all session in the institute and participants achieved the cut-off of 85% 239 correct responses after the final session (Fig 1 B). A final test on schema-knowledge (24 240 questions) about 14 days after the last session in the institute and directly after the 241 delayed retrieval of novel schema-related and control facts in the MR scanner (Fig 1B) 242 showed that participants had successfully acquired schema-knowledge. Participants 243 reached 91.8% (±1.4%) high confidence correct responses and across all confidence 244 levels 96.9% (±0.8%) correct responses (Fig 2B). The response times for high confidence 245 schema-knowledge retrieval was similar to high confidence immediate retrieval of novel 246 schema-related facts and substantially faster than their delayed high confidence retrieval, 247 which occurred in the same session (Fig. 2 D).

### 248 Learning exemplar names of the control schema

249 Differences in familiarity with the names of the schema and control exemplars could affect 250 memory formation independent of the hierarchical, associative knowledge structure that 251 was acquired only for the schema (Chua et al., 2012). The goal was therefore to minimize 252 potential differences in familiarity between the names of schema and control exemplars 253 prior to learning novel facts. Starting in session 2, participant were learning and writing 254 down the names of the control exemplars in random order without any information about 255 the nature of them. Participants expected to be tested on their memory for these 12 words 256 in the next session. Participants were asked to recapitulate the names of the control 257 exemplars in each of the following homework sessions and were tested for them in each of 258 the sessions in the institute. As intended, this procedure resulted in high familiarity and 259 perfect memory for the names of the exemplars of the control schema. However, this tight 260 experimental control also probably established to some degree semanticized and 261 arbitrarily interconnected representations of the control exemplar names, which potentially 262 reduces the observable prior knowledge effect.

### 263 Encoding and retrieval of novel schema-related and control facts

264 Encoding and retrieval of schema-related (SR) and not schema-related (NS) control facts 265 took place in the MR scanner. Three encoding rounds for SR and NS facts were followed 266 by four additional encoding rounds outside of the scanner only for NS facts to reach similar 267 performance in the following immediate memory test in the MR scanner. Similar memory in 268 the immediate retrieval test for novel schema-related and control facts is a critical 269 prerequisite to relate potential differences in forgetting until the delayed memory test to 270 divergent consolidation trajectories. About two weeks later (15.17 ±1.95 days) there was a 271 delayed retrieval test followed by the aforementioned retrieval of schema-knowledge.

272 The encoding and immediate retrieval rounds took place on the day following the 273 eighth session in the institute. After arrival, participants wrote down the names of the 12 274 schema and the 12 control exemplars in order to activate their schema-knowledge. 275 Participants were instructed to memorize each novel fact carefully, to focus only on the 276 facts that was presented at the time, and to give each facts equal memorization effort. 277 Participants were also informed about how they would be tested in the retrieval test. In 278 each of the three encoding rounds all novel 72 SR and 72 NS facts (6 for each exemplar) 279 were presented in pseudorandom order (consecutive facts were from different category 280 members). Each encoding round was distributed across two fMRI runs. Each trial started

281 with the presentation of a fact for 4s (Fig. 1 B). Participants were asked in the first round to 282 indicate within this time interval whether they will remember the fact (i.e. judgement of 283 learning; van Buuren et al., 2014; Witherby and Carpenter, 2021) and in the two 284 subsequent rounds whether they remembered the fact (judgement of memory). Therefore, 285 the tasks differed between the first and the later encoding rounds, which we considered 286 acceptable because implicitly there is always an unavoidable difference between the first 287 and subsequent encounters with new information. To foreshadow the results of our across 288 round representational similarity analyses (Figs. 5 and 6), similarity between the first two 289 rounds was rather greater which speaks against a major influence of the different tasks on 290 fMRI results. They had for each hand one button box, pressing the left index finger to 291 indicate 'yes, pressing the right for 'no'. The presentation of the facts was followed by a 292 fixation cross for 500ms, followed by an active baseline task. In this task the fixation cross 293 changed its color from white to either red or blue and participants were instructed to press 294 a corresponding button as quickly as possible. The baseline task was introduced to 295 prevent rehearsal of the previously encoded novel facts and to reduce activity in the 296 default mode network in the implicit baseline (Stark and Squire, 2001). This baseline 297 lasted between 2 - 4 s (jittered) before the next stimulus was presented. After the 3 298 encoding rounds, participants left the scanner and encoded the NS facts for 4 additional 299 rounds because we aimed for equal performance in the immediate memory test for NR 300 and NS facts. The previous study and further piloting showed that to achieve comparable 301 performance NS items need to be presented twice as many times as SR items (Hennies et 302 al., 2016). The procedure was the same as before.

303 Participants entered then the scanner again for the immediate retrieval test. All 72 304 SR and 72 NS facts were tested in a two-alternative forced choice task distributed across 305 two fMRI runs. Each trial started with the presentation of a fixation cross in the middle of 306 the screen between 2-4 sec (jittered). This was followed by the presentation of a fact 307 together with its alternative fact for 6 s. Participants had to choose the correct answer by 308 pressing the corresponding button on one of the two button boxes with the right or left 309 hand within the 6 s interval. Participants had to indicate their confidence by choosing 310 between the options sure, unsure, guess. After each trial the same active baseline task 311 that was employed during the encoding followed. Approximately 2 weeks later participants 312 returned for the delayed retrieval test in the scanner, which was identical to the first test, 313 but the order of facts was re-randomized. After retrieving all 72 SR and 72 NS facts 314 distributed across 2 fMRI runs participants were tested in a final fMRI run on 24 of

schema-knowledge facts that were tested in a similar format, i.e. forced choice withconfidence ratings.

317 It should be noted that the lure of a novel fact during retrieval (e.g. '4 cm' when 318 'Styga is 2 cm long' was learned) was not systematically associated as a novel fact with 319 another exemplar (e.g. '4 cm' was not necessarily a novel fact related to another 320 exemplar). This lack of a systematic using each novel fact as both, lure and target might 321 preclude an unambiguous interpretation of the results as noted by a reviewer: The novel 322 schema-related and control facts might have been more familiar than the lures during the 323 recognition tests and the responses could reflect familiarity driven item memory and not 324 assimilation into the schema. However, although we did not systematically used each lure 325 as target in a different recognition trial many lures or at least very similar responses 326 appeared in more than one trial which reduces this potential confound (a list of all retrieval 327 guestions and the response alternatives can be found at https://osf.io/aj28h/).

328 As described participants encoded and retrieved ('Did you remember?') the control 329 facts in seven rounds (compared to three rounds for schema-related facts) and 330 immediately retrieved them then. This many rounds was necessary to reach similar 331 performance in the immediate memory test, which was critical for the second goal of the 332 study, i.e. to test prior knowledge effects on consolidation. However, on the other hand the 333 repetitive encoding of object-location associations, respectively word lists has been 334 recently shown to result itself in accelerated system consolidation that is stabilized during 335 sleep (Brodt et al., 2016, 2018; Himmer et al., 2019). In the current study this effect might 336 be even stronger due to the over-learned and consolidated names of the control 337 exemplars. In other words, the tight experimental control might also have resulted in 338 consolidated memory traces for the novel control facts - even if via a different mechanism 339 as hypothesized for the novel schema-related facts.

### 340 fMRI data acquisition

Functional magnetic resonance imaging (fMRI) was performed on a 3 T system (Siemens
Trio) with a 32-channel head coil. An echo planar imaging T2\*-sensitive sequence in 50
contiguous axial slices (3 x 3 x 2.8 mm) with 1 mm gap; TR, 2.96 s; TE, 30 ms; PAT factor
2; flip angle, 80°; matrix 64 x 64) was employed. High resolution (1 x 1 x 1 mm voxel size)
T1-weighted structural MRIs were acquired for each subject using a 3D MPRAGE
sequence as part of the first scanning session.

### 347 fMRI data preprocessing

348 Functional imaging data were processed using the Statistical Parametric Mapping 12 349 software (SPM12, Wellcome Department of Cognitive Neurology, London, UK; 350 http://www.fil.ion.ucl.ac.uk/spm). Functional images were realigned and unwarped to 351 correct for susceptibility-by-movement artifacts. The anatomical image were coregistered to the mean functional image of that participant. The anatomical images were then 352 353 transformed into standard stereotaxic space using DARTEL as implemented in SPM12 354 and the deformation field applied to the functional images of the same participant. 355 Functional images were smoothed with full-width at half-maximum of 6 mm for the 356 univariate and of 3 mm for the multivariate analyses.

### 357 Univariate fMRI analyses

358 Individual subjects and group level data were analyzed using the general lineal model 359 (GLM) as implemented in SPM 12 in a mass univariate approach. One first level model was set up for encoding and retrieval. The two runs for each of the three encoding rounds 360 361 as well for the two runs of each immediate and delayed retrieval were concatenated where 362 the run specific constant, the autocorrelation structure and high pass filter were 363 appropriately adjusted. Regressors were created by convolving the onsets with the 364 canonical hrf. For each of the three encoding rounds one regressor for SR and NS facts 365 was created (due to ceiling effects we did not subdivide encoding events into those with a 366 positive and a negative judgment of learning, respectively memory). For immediate and 367 delayed retrieval, regressors were created for the high confidence correct, combined for 368 medium and low confidence correct and for incorrect responses as well as for the final 369 retrieval of the schema facts. In addition, six movement regressors were added as 370 nuisance variables.

371 On the second level, encoding related activity was analyzed by the main effect of 372 condition in an encoding round × condition (SR vs NS facts) ANOVA. The analysis of 373 retrieval activity was restricted to high confidence hits because for those a behavioral 374 effect of schema-knowledge was observed (Fig. 2 C) which is consistent with previous 375 literature (Long and Prat, 2002; Bein et al., 2020) and it has been argued high confident 376 responses are most informative (e.g., Xiao et al., 2016; Lee et al., 2019). Retrieval activity 377 was analyzed by the main effect of condition and the interaction of condition and delay in a 378 delay (immediate vs. delayed) × condition (SR vs NS facts) ANOVA. To identify areas that 379 might be involved in semantic memory, retrieval activity during high confidence retrieval of

the schema facts (after the delayed retrieval of novel facts) was contrasted against highconfidence immediate retrieval of novel schema-related facts in a paired t-test.

382 In addition, two Psycho-Physiological-Interactions (PPI; Friston et al., 1997) were 383 conducted based on previous literature (e.g., Sommer, 2017). The first PPI used as seed 384 the vmPFC cluster (thresholded p < 0.001 uncorrected) that was identified by the analysis 385 of encoding activity and contrasted its functional coupling during encoding SR and NS 386 facts for each of the three encoding rounds. On the second level, coupling differences 387 between conditions were analyzed across encoding rounds in an ANOVA with the factor 388 round. The second PPI used as seed the vmPFC cluster identified by the retrieval second 389 level model to be more active during retrieval of SR than control facts immediately and 390 delayed. This PPI contrasted coupling during immediate as well as delayed retrieval of SR 391 with control facts. On the second level, it was analyzed where this coupling increases from 392 immediate to delayed retrieval.

### 393 Multivariate fMRI analyses

394 In order to get parameter estimates for each individual trial as input for the multivariate 395 analyses, for each trial an independent first-level model was created with one regressor 396 containing only the corresponding trial and one for all other trials in that fMRI run (Mumford 397 et al., 2012). In addition, six movement regressors were added as nuisance variables as 398 well a high-pass filter applied and corrected for autocorrelation. The t-maps testing the 399 beta of the trial of interest in each model against the implicit baseline were used for the 400 following RSA to reduce the influence of noisy voxels (Dimsdale-Zucker and Ranganath, 401 2018). In all RSA we employed a whole brain searchlight approach (radius 3 voxels) and 402 correlated (Pearson's linear rank correlation) the resulting vectors of trial-specific t-values 403 across conditions of interest. The resulting correlation coefficients were averaged within 404 condition after Fisher's Z-transformation and saved as value for the center voxel of the 405 current searchlight.

### 406 Encoding-encoding similarity

In a first RSA we aimed to find support for a more rapid integration of novel schemarelated facts into the activated associative structure in terms of a greater consistency of
the distributed activity pattern across encoding rounds. Therefore we analyzed the
robustness of fact-specific activity patterns (e.g. *Texana is 2 cm long* or *NIV contains Copper*) between the succeeding encoding rounds, i.e. between round 1 and 2 as well as
between round 2 and 3 (Xue et al., 2010; Bruett et al., 2020). In particular the activity

pattern during encoding of a specific fact was correlated with encoding the same fact in the
succeeding round (similarity) as well as with encoding of all the other facts in that round
(dissimilarity), separately for novel schema-related and control facts. On the second level
we contrasted the similarity-dissimilarity difference maps in a 2 (round 1/2 vs. round 2/3) x
2 (SR vs NS) ANOVA.

### 418 Encoding-operation similarity analyses

419 In a second RSA we aimed to find support for more effective organization processing 420 during encoding of novel schema-related facts. Therefore, we aimed to identify brain 421 regions in which prior knowledge affects the encoding operation irrespective of the specific 422 to be encoded fact. To this end we correlated on the one hand activity during encoding 423 each novel schema-related fact with all other schema-related facts in a round, but 424 excluded the 5 other trials with facts related to the same exemplar (e.g. 'Texana') and on 425 the other hand activity during encoding of each novel schema-related fact with all control 426 trials resulting for each round in a SR-SR and a SR-NS correlation per subject. For the 427 latter we excluded also all facts related to one control exemplars to avoid different 428 numbers of correlations contributing to SR-SR and SR-NS similarities. On the second level 429 we contrasted the resulting similarity maps in a 3 (round 1 vs. 2 vs. 3) x 2 (SR-SR vs SR-430 NS) ANOVA.

To test whether prior knowledge also results in more similar activity patterns related to the encoding operation across rounds, we repeated the above analysis but correlated activity during encoding across succeeding round, e.g. each schema-related trial in round 1 with all other schema-related trials in round 2 except with the 6 trials related to the same exemplar (e.g. 'Texana'). On the second level we ran a 2 (round 1/2 vs. round 2/3) x 2 (SR-SR vs SR-NS) ANOVA.

In addition, with the same goal, we conducted two similar, complementary analyses
where we contrasted within and between rounds the correlation of activity during all novel
schema-related trials (but again excluded the trials related to the same exemplar) with the
correlation of activity during all control trials. This approach resulted at the second level in
a 3 (round 1 vs. 2 vs. 3) x 2 (SR-SR vs. NS-NS) and a 2 (rounds 1/2 vs. rounds 2/3) x 2
(SR-SR vs NS-NS) ANOVA.

443 Encoding-retrieval similarity

444 As an alternative approach to find support for differences in consolidation due to prior

445 knowledge we computed the similarity of activity patterns between encoding round 1 and

- 446 immediate as well as delayed retrieval for the novel schema-related and control facts. On
- 447 the second level we run a 2 (encoding-immediate vs. -delayed retrieval) x 2 (SR-SR vs.
- 448 NS-NS) ANOVA.
- Encoding schema knowledge retrieval similarity 449

450 To more directly test which areas are involved in the assimilation of the novel schema-451 related facts into the overlearned schemata, we correlated in another RSA activity during 452 encoding novel schema-related facts with activity during retrieval of the schema-453 knowledge (where we excluded again trials related to the same exemplar) and contrasted 454 it with the correlation of control facts with the retrieval of schema-knowledge. This results 455 at the second level in a 3 (round 1 vs. 2 vs. 3) x 2 (SR-Schema vs NS-Schema) ANOVA. 456 However, it should be noted that this analysis is to some extent confounded by the 457 shortcoming that the schema-knowledge was retrieved only after the delayed retrieval of 458 the novel schema-related and control facts. At this time point, the novel schema-related facts were at least partly already assimilated into the schema-knowledge which may have 459 460 led to changes in the representation of that initial schema itself, i.e. accommodation.

461 Multivariate–Univariate Dependence Analysis

462 Pattern similarity can be caused not only by distributed patterns of activity but also by 463 consistent (de-) activation of voxels in a univariate fashion. Therefore, we conducted a 464 multivariate-univariate dependence analysis (MUD) previously suggested by Aly and Turk-465 Browne (2015) for significant RSA-peak voxels that were in brain regions which also show 466 univariate effects. In particular, we first multiplied in each voxel of the spheres the 467 normalized values of the corresponding trials, e.g. encoding of a specific fact in round 1 468 and 2. These products indicate how much a voxel contributes to the correlation, i.e. to 469 pattern similarity. These products and the mean activity of each voxel were averaged 470 across trials and then correlated across voxels. The magnitude of the correlation indicates 471 how much univariate effects contribute to the RSA result.

472 Importantly however, a MUD correlation does not necessarily imply that the voxels 473 in a sphere show a univariate effect consistently in the same direction but only that voxels 474 activate or deactivate consistently for trials in a condition and that the same voxels 475 contribute to the observed similarity. Therefore, we computed in addition the correlation of 476 the differences in multivariate similarity between conditions in the RSA-peak voxels with 477 the difference in univariate mean activity in the searchlight-sphere around the RSA-peak 478 voxels across participants (Wagner et al., 2016). The differences to the MUD are that the

similarity difference in the RSA-peak voxel (i.e. the difference between the correlations
across all voxels in the surrounding searchlight sphere) and the difference in mean activity
in all voxels of the searchlight sphere are used. This approach is diagnostic of the extent to
which the observed similarity difference might be driven by differences in mean activity
between conditions.

### 484 Correction for multiple comparisons

485 Results of all fMRI analyses were considered significant at p < 0.05, family-wise-error 486 (FWE) corrected for multiple comparisons across the entire scan volume or within the a 487 priori defined anatomical regions of interest (ROIs). ROIs for the bilateral hippocampus, 488 bilateral precuneus/posterior cingulate and angular gyrus were computed from the 489 Harvard-Oxford cortical and subcortical structural atlases. A vmPFC ROI was manually 490 traced on the mean T1 image based on previously published post-mortem data (Mackey 491 and Petrides, 2014) using ITK-SNAP 3.6.0 (Yushkevich et al., 2006). The vIPFC ROI we 492 functionally defined as a sphere with radius 10 mm centered around the previously 493 observed peak voxel (xyz = [-40 4 28]; Sommer, 2017).

### 494 **Results**

### 495 Behavioral results

Encoding and retrieval performance was analyzed in mixed effects models using the R
base package and the *Ime4* as well as the *ImerTest* packages for computing type III
ANOVAs with Satterthwaite's method for the approximation to degrees of freedom (Bates
et al., 2015; Kuznetsova et al., 2017). Model fit, such as normality of model residuals, was
verified using the *check\_distribution* function in *R* package *performance* (Lüdecke et al.,
2020). Post-hoc Tukey HSD tests were performed using the *lemans* package (Lenth,
2016) if a paired comparison was of relevance for the interpretation of the results.

### 503 Encoding

504 Confirmative (i.e. 'yes') responses in the judgement of learning (round 1) and judgment of 505 memory (rounds 2 to 3) for schema-related and control facts were analyzed in mixed 506 effects models with the fixed effects condition (schema vs. control) and encoding round (1 507 vs. 2 vs. 3) as well as subject as a random factor. This model was compared with a similar 508 one but with a subject specific slope across rounds as additional random factor. Because 509 the model fit was similar ( $X^2(5) = 0.6$ , p = .99) we used the less complex model. The 510 effects of condition and round on judgments of learning, respectively memory (Fig. 2 A) 511 reached significance (F(1,135.00) = 376.35, p < 0.001; F(2,135) = 14.67, p < 0.001) but

not their interaction (F(2,135.00) = 0.49, p = 0.614). Participant's judgment of memory for 512 513 the control facts increased outside of the scanner (F(3,78) = 36.48, p < 0.001) where post-514 hoc Tukey HSD tests show that there was no increase between round 3 and 4 (p = 0.714) 515 but only between round 1 and 2 as well 2 and 3 (p < 0.001; p = 0.007) suggesting 516 participants performance reached an asymptote. A direct comparison with round 3 of 517 learning schema-related facts and round 7 of learning control facts showed less subjective 518 memory (t(26) = 2.48, p < 0.020) for control facts. 519 The effects of condition and round on reaction times (Fig. 2 B) also reached significance (F(1,81) = 16.813, p < 0.001; F(2,27) = 144.766, p < 0.001) as did their 520 521 interaction (F(2,81) = 10.005, p < 0.001). Restricting the analyses to confirmatory (i.e. 522 (yes') responses resulted in a similar pattern, i.e. significant main effects (F(1,81) = 9.532, 523 p = .003; F(2,27) = 186.87, p < .001) but no interaction (F(2,81) = 0.542, p = .584). 524 Reaction times did not further decrease significantly outside of the scanner during learning 525 round 4 to 7 for control facts (F(3,78) = 2.12, p = 0.104). Reaction times in the last round of 526 control facts encoding were faster than in the third and last round of schema-related facts

527 (t(26) = 2.61, p = 0.015).

### 528 Immediate and delayed retrieval

529 Hits during immediate and delayed retrieval were analyzed in mixed models with the 530 fixed effects condition (schema vs. control) and confidence (3 levels) as well as subjects 531 as a random factor with confidence as random slope (Fig. 2 C). This model had a 532 significantly better model fit than a less complex model without the random slope (immediate retrieval  $X^{2}(5) = 174.1$ , p < .001; delayed retrieval  $X^{2}(5) = 147.8$ , p < .001). 533 534 Immediate retrieval showed only a significant effect of confidence level (Fig 2 B; F(2,27) =535 264.33, p < 0.001, but not of condition, and no interaction (F(1,108) = 0.0037, p = 0.951; F(2,108) = 0.076, p = 0.923). In other words, the additional 4 encoding rounds outside of 536 537 the scanner for control facts resulted as intended in a similar immediate hit rate for 538 schema-related and control facts. Therefore, potential difference in delayed memory 539 performance cannot be attributed to differences in the initial memory strength. For delayed 540 retrieval, there was again a significant effect of confidence level (F(2,27) = 24.05, p < 541 0.001), and also a significant interaction (F(2,81) = 7.13, p = 0.001) which was driven by 542 more high confidence hits for schema-related than control facts (post hoc Tukey HSD p = 543 0.014), but no effect of condition (F(1,81) = 0.32, p = 0.585). In a mixed model including 544 both retrieval tests with the additional factor and random slope delay (immediate vs.

545 delayed) the interaction of condition, delay and confidence reached only a trend towards 546 significance (F(2,269.99) = 2.63, p = 0.074) besides the significant effects of delay,

- 547 confidence and their interaction. (Loftus and Masson, 1994)
- 548

549 The proportion of incorrect responses across the confidence levels was also 550 analyzed in order to test whether prior-knowledge might result in relative higher proportion 551 of high confidence incorrect responses. Neither during immediate nor delayed retrieval 552 was the interaction of schema and confidence significant (immediate retrieval: effect of 553 schema F(1,168) < 0.01, p = .971; effect of confidence F(2,168) = 324.26, p < .001; 554 interaction  $F(2,168) = \langle .01, p = .991 \rangle$ ; delayed retrieval: effect of schema F(1,168) = 0.03. 555 p = .872; effect of confidence F(2,168) = 31.03, p < .001; interaction F(2,168) = 1.28, p 556 = .28).

557 Reaction times for hits were analyzed in similar mixed models (Fig. 2 D). For 558 immediate retrieval both main effects as well as the interaction reached significance 559 (F(1,101.34) = 55.61, p < 0.001; F(2,38.19) = 40.30, p < 0.001; F(2,101.04) = 31.68, p < 0.0560 0.001) indicating overall slower retrieval of schema-related facts, where there was no 561 difference for high confidence but for lower confidence hits. Also, for delayed retrieval both 562 main effects and the interaction reached significance (F(1,100.96) = 8.49, p = 0.0041; 563 F(2,47.95) = 47.95, p<0.001; F(2,100.90) = 3.68, p = 0.029) suggesting a similar pattern. 564 i.e. no difference for high confidence hits. In a mixed model including both retrieval tests 565 with the additional factor and random slope delay, the effects of condition (F(1,225.86) =566 43.13, p < 0.001), confidence (F(2,37.08) = 50.45, p < 0.001), and delay (F(1,28.02) = 567 59.27, p < 0.001), as well as the condition × delay (F(2,226.13) = 20.17, p < 0.001) and the 568 confidence × delay interaction (F(2,225.33) = 4.35,p = 0.014) reached significance 569 indicating overall slower retrieval and a smaller difference between conditions after the 570 delay.

### 571 Univariate fMRI results

### 572 Encoding

At first, we present activity related to retrieval of the overlearned schema-knowledge although it was assessed last (i.e. after delayed retrieval of novel schema-related and control facts) because its consolidation is the basis for the assimilation of novel schemarelated facts. We contrasted retrieval of schema-knowledge with immediate retrieval of novel schema-related facts because both refer to schema-exemplars and were similarly 578 fast. Activity during retrieval of the over-learned schema-knowledge was greater in the 579 vIPFC and other areas (Fig. 3 A, Table 1). Activity in the other three conditions, i.e. 580 immediate retrieval of control facts as well as delayed retrieval of schema-related and 581 control facts, is plotted in transparent bars because these conditions did not contribute to 582 the statistical test that identified this area. We present it to show that activity did not differ 583 between the immediate and delayed retrieval of novel-schema-related and control facts. 584 This plot shows that activity during retrieval of the schema-knowledge was also greater 585 than during immediate retrieval of control facts and delayed retrieval of novel schema-586 related and control facts.

587 During encoding the main effect of SR > NS across rounds was significant in the 588 vmPFC, in a cluster comprising the ventral precuneus/retrosplenial cortex (vPC/RSC) as 589 well as in the superior parietal cortex (Fig. 3B) implicating greater activity during encoding 590 of novel schema-related facts. The interaction of condition and round showed that in the 591 vmPFC and vPC/RSC the difference was greater in the first than the last round (Table 1 for 592 full list of results). Moreover, the vmPFC was more strongly coupled with the hippocampus, 593 fusiform, supramarginal and inferior frontal gyri, dorsal precuneus and superior parietal 594 cortex during encoding of schema-related than control facts (Fig. 3 C). The interaction with 595 round showed that this difference in coupling in the hippocampus and dorsal precuneus 596 was largest in round 1.

### 597 Retrieval

598 The vmPFC and the vPC/RSC were also more active during retrieval of novel schema-599 related than control facts irrespective of delay (Fig. 4 A). The PPI using the vmPFC cluster 600 as seed revealed stronger coupling differences with the precuneus/posterior cingulate 601 between retrieval of SR and NS facts after the delay (Fig. 4 B). A region of the vmPFC, in 602 particular the subgenual ACC, showed an increase in activity only during delayed retrieval 603 of schema-related facts whereas the hippocampus revealed a decrease in activity from 604 early to delayed retrieval only of control facts (Fig. 4 C). There was no such interaction 605 effect in the vIPFC ROI (largest Z = 1.43, p = 0.661 at [-42 0 18]) contrary to our previous 606 study (Sommer, 2017).

### 607 Multivariate fMRI results

### 608 Encoding-encoding similarity

609 In Fig 5 A right panel we present the results of the encoding-encoding similarity analysis

610 irrespective of prior knowledge (i.e. the main effect similar greater dissimilar) in order to

611 visualize its sensitivity because contrasting schema-related and control facts using this 612 measure revealed only relatively subtle differences. In particular, pattern robustness in 613 terms of encoding-encoding similarity between rounds was greater for novel schema-614 related than control facts in the right inferior frontal gyrus (IFG, [48 12 21], Z = 3.90, Fig. 5 615 A left panel). Importantly though, as the IFG was not an a priori defined ROI the peak 616 would not survive correction for multiple comparisons. We decided to still report and 617 visualize it for exploratory reasons because the IFG has been observed before to be more 618 active and stronger coupled with the hippocampus during schema retrieval (Bein et al., 619 2014; van Buuren et al., 2014; Wagner et al., 2015).

620 The interaction of similarity and encoding round showed that encoding-encoding 621 similarity between the first two rounds was greater for schema related items in early visual 622 cortex ([0 -90 3] Z = 3.93, Fig. 5 B left panel) and the amygdala ([27 -3 -18], Z = 4.10) and 623 between round 2 and 3 in the precuneus ([9 -72 39] Z = 4.78, Fig 5B right panel) where 624 only the latter peak reached significance corrected for multiple comparisons. Because the 625 precuneus also showed univariate effects (Fig 3 A) we conducted a multivariate-univariate 626 dependence analysis (MUD; see methods; Aly and Turk-Browne, 2015) which revealed no 627 correlation between the observed multivariate effect and the univariate effect (r = -.098, p 628 = .122).

### 629 Encoding operation similarity analysis

630 The RSA analysis comparing correlations between encoding of schema related facts with 631 schema related facts (SR-SR) and the correlation between encoding schema related facts 632 and novel facts (SR-NS) within each round revealed the vmPFC ([3 51 -9], Z = 5.66) and vPC/RSC ([-15 60 21], Z = 6.72) as well as the midcingulate gyrus ([0 -15 48], Z = 5.23), 633 634 left middle frontal gyrus ([-21 15 42], Z = 4.82) and the right central operculum ([48 -18 21], 635 Z = 4,57) as a main effect across the three rounds. Notably, when multiple facts related to 636 the same exemplar they were excluded. Because the first two areas also showed 637 univariate effects we conducted MUD analyses for the peak voxels which revealed no 638 correlation for the vmPFC (r = .028, p = .694) but did show a weak correlation for the 639 vPV/RSC (r = .254, p = .006). Therefore, we correlated in addition the differences in 640 multivariate similarity in the RSA-peak voxels and the differences in univariate mean 641 activity in the searchlight-spheres around the RSA-peak voxels between conditions across 642 participants which revealed no relationship (r = .301, p = .115; r = .189, p = .337)

643 suggesting no major contribution of univariate activity differences between conditions to644 the multivariate results.

645 The interaction between the within-round similarity and round (Fig. 6 A) showed 646 greater similarity in the vmPFC ([-3 48 -6], 6.37), vPC/RSC ([-9 -60 30], Z = 5.45), bilateral 647 angular gyrus ([-48 -60 24], Z = 4.56; [51 -51 27], Z = 4.07), OFC ([-27 36 -12], Z = 7.52) 648 and left hippocampus ([-33 -24 -12], Z = 3.75) in the first compared to the third round. For 649 the first two peaks we again computed a MUD analysis to disentangle uni- and multivariate 650 contributions. This showed subtle but significant correlations between uni- and multivariate 651 effects (vmPFC: r = .113, p = .038; vPC/RSC: r = .249, p < .001). Therefore, we correlated 652 again in addition the multivariate difference in the peak-voxels and the mean univariate 653 differences in the corresponding searchlight sphere across participants which revealed 654 only a trend towards significance for the vPC/RSC (r = -.225, p = .202; r = .323, p = .094).

655 The across round encoding operation similarity analysis (Fig. 6 B) revealed as main 656 effect (i.e. across rounds) the vmPFC (-9 51 0], Z = 3.50) and the vPC/RSC ([-12 -57 15], 657 Z = 4.87; [15 -54 18], Z = 4.28) but no interaction with round. The MUD analyses showed a 658 subtle but significant correlation only in the vmPFC (r = .244, p < .001; r = -.056, p = .495; r659 = .028, p = .686) where the additional follow-up correlation of RSA-peak differences in 660 similarity and mean activity in the corresponding searchlight-spheres suggested that the 661 univariate contributed r = -197, p = .316; r = .280, p = .150; r = .442, p = .019) only to the 662 second peak in the vPC/RSC.

663 In addition we computed a complementary within-round encoding operation 664 similarity analysis in which we contrasted the correlation between the novel schema-665 related facts (SR-SR) with the correlation of the control facts (NS-NS), excluding the trials 666 related to the same exemplar. This also revealed greater similarity across rounds in the 667 vmPFC ([-12 57 -3], Z = 5.79) and precuneus/RSC ([-15 -60 21], Z = 4.59; [-6 -48 12], Z = 668 6.13). Because both areas also showed univariate effects we conducted a multivariate-669 univariate dependence analyses (Aly and Turk-Browne, 2015) for the three peak voxels. 670 The univariate contribution to the observed similarity differences correlated towards a 671 trend in the vmPFC (r = .105, p = .085), and in the first peak in the vPC/RSC (r = .003, p 672 = .096; r = -.037, p = .574). Thus, in the vmPFC and parts of the vPC/RSC univariate 673 effects might have contributed modestly to the similarity.

674The interaction of round and schema-related vs. control showed that the difference675in similarity was greater in the first round in vmPFC (Fig. 6 A; [-9 57 0], Z = 5.98), OFC ([-

676	27 36 -12], Z = 7.25), bilateral angular gyrus (Fig 6 A; [-48 -66 21], Z = 4.18; [51 -51 27], Z
677	= 4.13), and the vPC/RSC ([-3 -54 18], Z = 4.10). The multi-univariate dependence
678	analyses for these peaks revealed a marginal trend towards a correlation only for the
679	precuneus (r = .092, p = .089). Therefore, pattern similarity in the precuneus appears to
680	be, a least to some degree, related to unidirectional changes in voxel activity. Across
681	rounds similarity of encoding operation was also greater for schema than control trials in
682	the vmPFC (Fig. 6 B; [-9 54 0], Z = 4.37) and vPC/RSC ([-6 -48 12], Z = 6.30; [6 -45 15], Z
683	= 5.20). The univariate-multivariate dependence analyses revealed no significant
684	correlation (r = .083, p = .130; r = .050; p = 380, r = .008, p = .859).

685 Encoding-retrieval similarity

The contrast of encoding-retrieval similarities showed higher similarity during encoding and retrieval of novel schema-related facts at both delays in the middle temporal ([54 -57 3], Z = 5.31) and left angular gyrus ([-45 -57 42], Z = 4.23). The interaction revealed an increase in the parahippocampus ([27 -36 -15], Z = 4.08).

690 Encoding - schema knowledge retrieval similarity

691 We observed a main effect of greater similarity between encoding novel schema-related 692 than control facts with the retrieval of schema-knowledge in the vmPFC ([-9 51 0], Z = 693 5.36, Fig. 6 C), the vPC/RSC(-3 -54 18], Z = 4.51; [-15 -60 21], Z = 3.78) and the posterior 694 cingulate ([0.30 36], Z = 5.34). The MUD analyses for the regions where we observed 695 univariate effects showed subtle but significant correlations for the vmPFC and one peak 696 in the vPC/RSC (r = .215, p = .001; r = .150; p = .008; r = .053 p = .400). However, the 697 individual differences in multivariate similarity and univariate mean activity did not correlate 698 across participants (r = -.103, p = .602; r = .065, p = .743; r = .046, p = .815) suggesting 699 that the similarity difference was not caused by a consistently greater activity in the voxel 700 of the sphere.

Finally, an interaction with round in terms of greater difference in round 1 than round 3 was observed in the posterior cingulate ([-3 -30 36], Z = 4.64), the left middle frontal gyrus ([-27 24 45], Z = 5.52), right middle temporal gyrus ([57 -21 -18], Z = 5.09), as well as trends towards significance in the bilateral angular gyrus ([-42 -60 21], Z = 4.55; [51 -33 33], Z = 4.46).

### 706 Discussion

707 The over-learning procedure in the current study likely resulted in semanticized, cortical 708 representations of the schema-knowledge. Learning of novel schema-related but 709 expectation-neutral facts strongly benefited from this prior knowledge. On the neural level, 710 we observed enhanced vmPFC-hippocampal coupling when information can be 711 assimilated in prior-knowledge. Not only was mean activity greater in vmPFC and the 712 vPC/RSC but also the distributed activity patterns in these areas showed greater similarity, 713 i.e. consistency for schema-based encoding operations with the angular gyrus within and 714 between rounds. Consolidation of the assimilated facts was also enhanced, as reflected by 715 slightly higher confidence retrieval and an increase in vmPFC activity and vmPFC-716 precuneus coupling.

### 717 Acquisition of schema-knowledge

718 The repetitive reactivation of the hierarchical schema-knowledge in various contexts (in the 719 institute, at home, on computer screens, as written handouts) and retrieval formats (essay-720 like texts, free recall, multiple-choice questions, pictures) across 7 weeks was expected to 721 result in semanticized and consolidated associative knowledge structures (Sekeres et al., 722 2018; Ferreira et al., 2019). The close to ceiling performance already early in learning (Fig 723 1 B) and, even more so, in the schema-knowledge memory test 14 days after the last 724 training session (Fig. 2 C) where response times were very fast (Fig. 2 D) show that the 725 knowledge was highly overlearned and likely semanticized. The strong involvement of the 726 vIPFC in retrieval of schema-knowledge further supports its proposed semanticization

- 727 because this area has been implicated in semantic memory and in retrieval of
- semanticized memory (Binder and Desai, 2011; Sommer, 2017). Taken together, the
- 729 acquired schema-knowledge very likely fulfilled the previously schema-defining criteria of
- 730 being an associative network structure, based on multiple episodes, lacking unit detail,
- 731 being adaptable as well as being cortical, i.e. semantic, representations (van Kesteren et
- 732 al., 2012; Ghosh and Gilboa, 2014).

### 733 Encoding of novel schema-related facts

- 734 The substantially higher judgments of memory for novel schema-related facts together with
- the more learning rounds for control facts that were necessary to reach similar
- 736 performance in the immediate memory test illustrate the power of the prior-knowledge
- 737 effect in the current paradigm (Fig. 2 A). Due to the invented schemata we can rule out the
- 738 possibility that congruency with schema-driven expectations or differences in
- 739 interest/motivation to learn the novel facts had a systematic effect on this mnestic

740 advantage (Chua et al., 2012; Witherby and Carpenter, 2021). The possibility that the 741 much more rapid learning of novel schema-related facts was not driven by their 742 assimilation but rather based on better item-level memory, i.e. familiarity based 743 recognition, cannot be ruled out. However, the only difference between retrieval of novel 744 schema-related and control facts was observed in terms of more high confidence 745 responses which are unlikely driven by familiarity and schema-knowledge is known to 746 impact predominantly recollection-based recognition (Long and Prat, 2002; Brandt et al., 747 2005). Taken together, the observed effect is likely driven by more effective organizational 748 processing which results in assimilation of the novel facts into the pre-activated associative 749 network (Ericsson and Kintsch, 1995) but schema effects on item-memory cannot be ruled 750 out.

On the neural level, the much more efficient encoding of novel schema-related facts 751 752 was paralleled by higher coupling of the vmPFC and hippocampus in addition to higher 753 mean activity in the vmPFC and the vPC/RSC where in the hippocampus no activity 754 difference was observed (Fig. 3 B/C). These results conceptually replicate our previous 755 findings using very different but also experimental schemata, i.e. associative structures of 756 random object-location associations, that also did not result in expectations about the 757 nature of novel facts, and also another study employing arbitrary expectation-neutral 758 information (Liu et al., 2016; Sommer, 2017). Together, these studies therefore support a 759 model that predicts stronger vmPFC-hippocampal coupling when novel expectation-neutral 760 information can be related to prior-knowledge (Gilboa and Marlatte, 2017). On the first 761 sight this interpretation stands in contrast to the prominent SLIMM model that proposes 762 that the vmPFC inhibits hippocampal encoding when it detects information congruent to 763 our expectations, i.e. reduced vmPFC-hippocampus coupling and less hippocampal 764 activity when schema-congruent information is encoded (van Kesteren et al., 2012; Greve 765 et al., 2019).

766 However, two distinct prior-knowledge effects might exist that are mediated by 767 qualitatively different encoding operations: When schemata allow specific expectations 768 based on previous encounters with the to be encoded information (i.e. palm-trees at a 769 beach) there is reduced vmPFC-hippocampus coupling when congruent information is 770 encoded (van Kesteren et al., 2012). This type of prior-knowledge effects results in 771 enhanced gist memory prone to schema-based distortions. However, when expectation-772 neutral novel information can be assimilated into a schema, e.g. a novel fact in our 773 academic discipline, that is further developed by this (i.e. accommodation) vmPFC-

hippocampal coupling is increased which results in enhanced and more accurate memory
formation (Long and Prat, 2002; Brandt et al., 2005; Tse et al., 2007).

776 The multivariate analyses show that more efficient encoding of novel schema-777 related facts seems not to be predominantly reflected in less encoding variability perhaps 778 due to the rapidity of integration (Fig. 5). The greater item-specific similarity in the IFG for 779 novel schema-related facts - although not predicted and hence not significant after 780 correction for multiple comparisons – would be consistent with this structure's previously 781 described role in representing dissimilarities during knowledge integration (Schlichting et 782 al., 2015). However, prior-knowledge substantially enhanced the consistency of the 783 encoding operations in the vmPFC, vPC/RSC and angular gyrus suggesting more effective 784 organizational processing (Fig. 6). In the first two areas we observed higher mean activity 785 replicating earlier findings with expectation-neutral (Maguire et al., 1999; Tse et al., 2011; 786 van Kesteren et al., 2014; Liu et al., 2016; Sommer, 2017) but also schema-congruent 787 facts (van Kesteren et al., 2013; Bonasia et al., 2018). The multivariate-univariate 788 dependence analyses suggested that the two effects might reflect different processes or 789 distinct aspects of the same process. Activity in the vPC/RSC has been associated with 790 retrieval, in particular with retrieval during encoding of novel information (Huijbers et al., 791 2012; Sestieri et al., 2017; van Kesteren et al., 2020) - consistent with activation of 792 schema-knowledge during encoding. The greater consistency of the distributed activity 793 patterns across all schema-trials suggest that irrespective of the current to be encoded fact 794 (e.g. 'Texana is 2 cm long') the superordinate schema-knowledge is activated.

795 The greater similarity of activity patterns during encoding of novel schema-related 796 than control facts with the retrieval of the overlearned schema-knowledge supports this 797 interpretation. However, because the schema-knowledge was only retrieved after the 798 potential assimilation of the novel information might have been already complete, this 799 similarity might reflect both, effects of assimilation and accommodation. This interpretation 800 would also apply to the greater consistency of encoding operations in the angular gyrus. 801 Interestingly, the angular gyrus has been - also by using multivariate analyses of activity 802 patterns - implicated in combining different schema components when it is applied to novel 803 related information (Wagner et al., 2015). The vmPFC has been suggested to bind 804 together co-activated vPC/RSC and angular gyrus representations to form a superordinate 805 knowledge template and to maintain the active schema when novel information is 806 processed (Gilboa and Marlatte, 2017). This interpretation of our multi- and univariate 807 results would be consistent with the more effective organizational processing of the novel

related information proposed by cognitive psychologists (Ericsson and Kintsch, 1995), i.e.
the integration in the associative network which leads to referential connections and the
association with appropriate retrieval cues.

The difference between encoding of schema-related and control facts was larger early in learning in several analyses. It is possible that overlearning the names of the control exemplars resulted in an arbitrary associative structure of these meaningless terms. After participants associated facts with the elements of this structure in the first round, it became to some extent schema-like. Alternatively, after the first round the novel schema-related facts may have already been assimilated to a large degree, which would also reduce the difference to encoding of the control facts.

### 818 Consolidation

819 The phenomenon that prior-knowledge leads also to more efficient consolidation has been 820 described only relatively recently (compared to the long history of schema-effects in 821 memory) in the aforementioned animal studies using novel expectation-neutral schema-822 related information (Tse et al., 2007, 2011). We confirmed this effects in humans in our 823 previous study where we translated the animal experiment to an fMRI design (Sommer, 824 2017). In the current study, using very different experimental schemata, we conceptually 825 replicated this effect. The impact on consolidation was rather subtle and was specific to 826 high confidence responses which is however consistent with previous reports on prior-827 knowledge effects in recognition memory (Long and Prat, 2002; Brandt et al., 2005). The 828 subtly of the prior-knowledge effect on consolidation might be also a side effect of the 829 seven learning rounds needed for the control information to reach similar immediate 830 memory performance because repetitive encoding and retrieval might result itself in faster 831 systems consolidation – although via different mechanisms (Brodt et al., 2016, 2018; 832 Himmer et al., 2019). In the current study this effect might be even stronger due to the 833 over-learned and consolidated names of the control exemplars. Using the same paradigm, 834 we previously showed that sleep spindle density as a proxy for nightly replay and systems 835 consolidation predicted the individual benefit of prior-knowledge on novel related facts 836 (Hennies et al., 2016). This finding supports the interpretation that the reduced forgetting 837 we observed in the current study is caused by more efficient systems consolidation. 838 Prior-knowledge resulted in an increase in vmPFC-precuneus coupling and of

wmPFC and hippocampus activity from immediate to retrieval of schema-related facts two
weeks later. In our previous study we did not observe the vmPFC but vIPFC, i.e. a
semantic memory area, to contribute more strongly to the retrieval of assimilated

842 information after two weeks (Sommer, 2017). This difference might be caused by 843 differences in the experimental designs, for instance that the schemata in our previous 844 study were much simpler, no meaningful hierarchical associative structures and probably 845 semanticized to a larger degree by even more intense over-learning. However, the vmPFC 846 (and RSC) has been implicated before in the retrieval of assimilated schema-related facts 847 (Tse et al., 2011) and the parallel relative increase in hippocampal activity would be 848 consistent with the incidental retrieval of associated schema knowledge during high 849 confidence recognition (Schultz et al., 2022). This parallel involvement of cortical, i.e. 850 vmPFC and RSC, and hippocampal retrieval would be consistent with the Trace 851 Transformation theory that proposes that both episodic and cortical traces can exist in 852 parallel (Sekeres et al., 2018). 853 Conclusions

854 The increased vmPFC-hippocampal coupling during the highly efficient encoding – likely 855 due to their assimilation - of novel schema-related expectation-neutral facts suggests a 856 prior-knowledge effect which is distinct from situations where the prior-knowledge allows 857 expectations. Together, our uni- and multivariate results support cognitive and 858 neuroscientific models about the processes underlying the putative assimilation, i.e. that a 859 vmPFC, vPC/RSC, angular-network results in the activation of schema knowledge 860 enabling more effective organizational processing of novel related facts. Moreover, the 861 results confirm that assimilation of novel related information also results in more effective 862 consolidation which is reflected, for not fully semanticized information, in vmPFC activity. 863

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### 993 Table 1: univariate fMRI results

contrast	area	xyz coordinate peak voxel	Z-value peak voxe
	encoding	·	
	vmPFC	0 54 -15	4.34
	ventral precuneus/RSC	-9 -57 15	6.36
SR > NS		12 -57 18	4.67
main effect	superior parietal cortex	-36 -78 39	5.34
		48 -72 33	4.79*
	post cingulate cortex	-3 -36 33	4,81
	frontal pole	-30 51 21	5.00
		36 54 18	5.32
NS > SR	anterior cingulate cortex	6 36 18	5.69
main effect	occipital pole	9 -81 -3	6.94
	lateral occipital cortex	-45 -78 0	4.85
		33 -90 -9	4.78
	supramarginal gyrus	57 -39 24	4.95
SR > NS	inferior frontal gyrus/insula	-51 -6 0	4.82
increase across rounds		54 12 -3	4.88
	anterior cingulate cortex	-3 33 15	4.93
	vmPFC	-3 51 -32	4.70
	ventral precuneus/RSC	-12 -57 15	4.09
SR > NS	dorsal medial PFC	-3 36 39	6.07
decrease across rounds	inferior frontal gyrus	-45 39 -12	5.92
	dorsolateral PFC	-42 15 42	4.88
	hippocampus	-21 -24 -15	3.99
	fusiform gyrus	-30 -57 -15	5.33
		33 -54 -12	6.00
	inferior frontal gyrus	48 36 3	5.63
PPI SR > NS	superior parietal cortex	15 -66 51	5.04
seed vmPFC		33 -42 42	4.91
main effect	dorsal precuneus	-9 -39 57	5.03
		-9 -69 51	4.89
	supramarginal gyrus	-57 -45 27	4.98
PPI SR > NS	hippocampus	-27 -21 -21	3.63
seed vmPFC		15 -12 -21	5.09
decrease across rounds	dorsal precuneus	9 -63 51	6.69
	retrieval		
	vmPFC	3 30 -21	4.57
SR > NS	ventral precuneus/RSC	-9 -57 15	6.49
		9 -54 15	5.68
	anterior cingulate cortex	-6 33 21	5.06
	insula	-39 15 -9	4.97
NS > SR		33 18 -15	4.46*
	frontal pole	24 57 -3	4.66
increase SR > NS	vmPFC	-6 18 -9	3.76

	hippocampus	27 -24 -15	3.74
PPI increase SR > NR seed vmPFC	ventral precuneus	15 -48 33	4.23
	schema-knowledge		
	ventro-lateral PFC	-42 9 30	3.99
	anterior cingulate cortex	9 18 39	5.31
schema retrieval >	ventral precuneus	21 -66 3	5.04
immediate novel SR	insula	30 24 -9	5.09
retrieval		-30 24 -6	4.96
	ventral striatum	-18 9 -6	4.84
		18 25 -6	5.58

994

995 Peal coordinates in MNI space. Correction for multiple comparisons was done on the

996 whole-brain level or within pre-defined anatomical regions of interest, specifically the

997 hippocampus, precuneus/retrosplenial cortex (RSC) and ventromedial prefrontal cortex

998 (vmPFC). \* trend towards significance

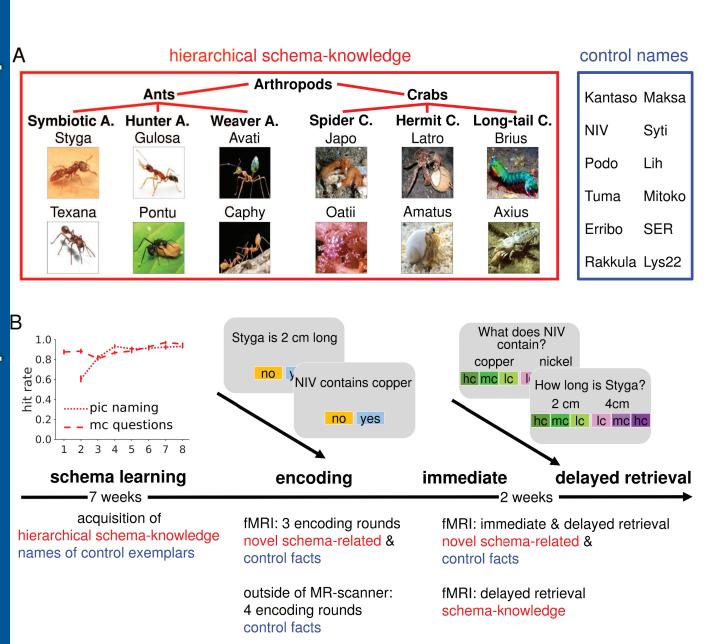
Figure 1. Schema and timeline of the experiment. A) Hierarchical structure of one of the two schemata (arthropods) and exemplar names of the other schema (cells) which served as control in this example. Schema and control were randomized across participants. See Hennies et al., 2016 for a figure with the hierarchical structure of the 'cell'-schema. B) Acquisition of schema knowledge and familiarization with the control names over 7 weeks with 1 learning session per week in the institute and homework in-between. Participants achieved high performance in the multiple-choice questions (mc questions) and the picture naming task of their schema. In the scanner, participants encoded 3 times (encoding round 1 to 3) 72 novel facts related to the exemplars of their schema and 72 facts related to their control exemplars. In encoding round 4 to 7 outside of the scanner, they only repeated the control facts to ensure equal immediate memory for schema-related and control facts In the first encoding round, participants judged whether they will remember the novel facts in rounds 2 to 7 whether they did remember it. Encoding was followed by immediate retrieval of all learned novel facts in the scanner. 2 weeks later all facts were retrieved again in the scanner followed by retrieval of 24 of the overlearned schema-knowledge facts During retrieval two equally plausible response alternatives were presented (targets and lures were randomized across participants) and participants indicated their confidence on a 3-point scale (hc – high, mc – medium, lc - low confidence).
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1017 Figure 2. Behavioral results. A) Encoding rounds 1 to 3 for novel schema-related and control facts took 1018 place in the MR scanner, rounds 4 to 7 only for control facts outside of the scanner. In the first round partici-1019 pants rated whether they will remember the fact (judgment of learning) and in round 2 to 7 whether they did 1020 remember the fact (judgement of memory). B) Response times for the judgment of learning (round 1), re-1021 spectively judgment of memory (round 2 to 7) during encoding. C) Proportion of high (hc), medium (mc) and 1022 1023 low confident (Ic) hits (relative to all responses in that delay x schema condition) during immediate and delayed retrieval for the schema-related and control facts as well as for the a subset of facts of the schema 1024 knowledge only during delayed retrieval. D) Retrieval times for high, medium and low confident hits during 1025 retrieval. Error bars are standard errors of the mean around the mean, corrected for interindividual differ-1026 ences (Loftus and Mason 1994). 1027

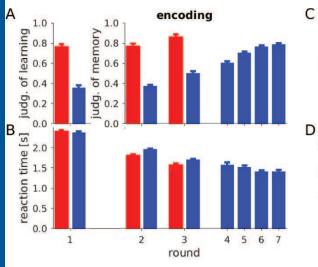
1043Figure 4. Activity differences during retrieval. A) During immediate and delayed retrieval of schema-related<br/>(SR, red bars) facts activity was greater in the vmPFC and the vPC/RSC. B) The Difference in coupling of<br/>the vmPFC with the precuneus during retrieval of schema-related (SR) and control (NS) facts increased from<br/>immediate to delayed retrieval. C) The vmPFC (subgenual ACC) and the hippocampus showed a larger<br/>activity increase from immediate to delayed retrieval of schema-related (SR, red bars) than control (NS, blue<br/>bars) facts. Error bars are standard errors of the mean around the mean, corrected for interindividual<br/>differences (Loftus and Mason 1994).Visualization threshold p < 0.001,uncorrected.</th>

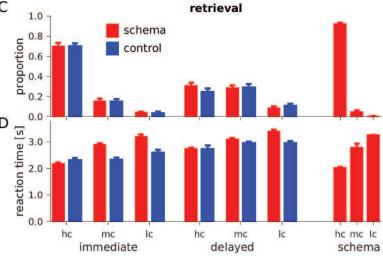
1052succeeding rounds was1053the right IFG (left panel)1054of the main effect. B) If1055cortex (left panel) and1056than control facts (NS1057comparisons and are	coding similarity (pattern robustness). A) Encoding-encoding similarity between s greater for novel schema-related (SR, red bars) than control facts (NS, blue bars) in I). The overall sensitivity of this approach is visualized in the right panel of A in terms incoding-encoding similarity between the first two rounds was greater in early visual between round 2 and 3 in the precuneus (left panel) for novel schema-related (SR). Note that the IFG and cuneus cluster are not significant corrected for multiple eported for exploratory reasons. Error bars are standard errors of the mean around r interindividual differences (Loftus and Mason 1994). Visualization threshold p <
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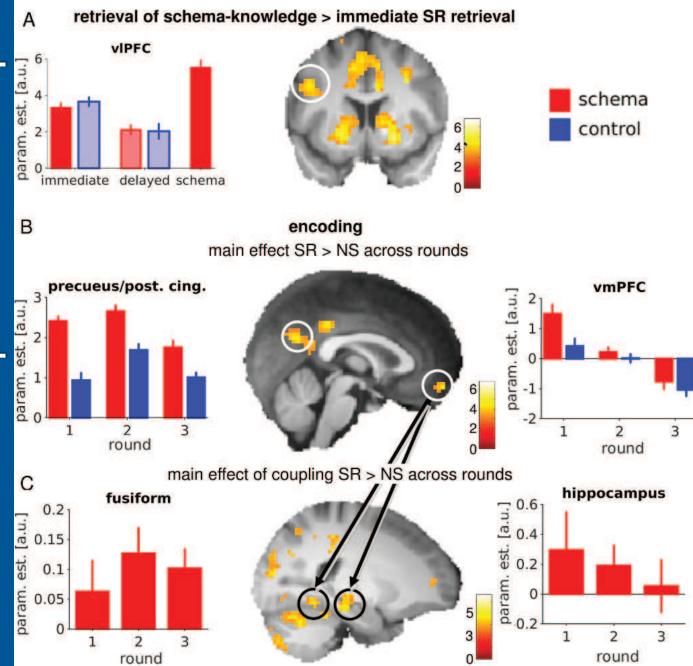
1061 Figure 6. Encoding operation similarity. A) Encoding-operation similarity within rounds was early in learning 1062 greater between encoding of novel schema-related (SR-SR, red bars) than between schema-related and 1063 control (SR-NS, blue bars) facts in the vPC/RSC, vmPFC, and the bilateral angular gyrus. B) Encoding-1064 operation similarity between rounds during encoding of novel-related (SR-SR) facts was also grater 1065 compared to SR-NS facts in the vPC/RSC and vmPFC. C) Operation similarity between encoding of 1066 schema-related facts and retrieval of schema knowledge was also greater compared to the encoding of 1067 control facts in the vPC/RSC and vmPFC. Error bars are standard errors of the mean around the mean, 1068 corrected for interindividual differences (Loftus and Mason 1994). Visualization threshold p < 0.001, 1069 uncorrected.

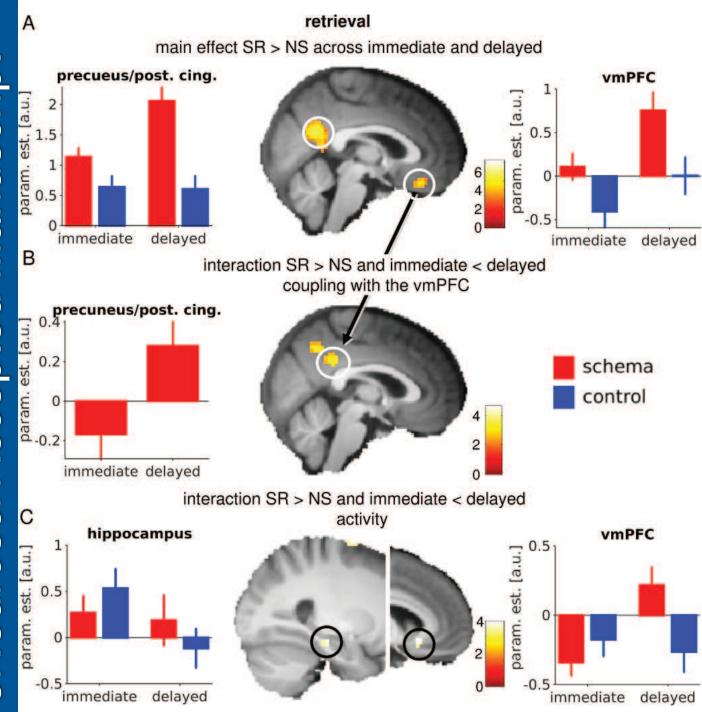


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