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### Implantable CMOS Pixel Sensor for Positron Imaging in Rat Brain

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#### Abstract

IMIC is a Monolithic Active Pixel Sensor prototype designed for the MAPSSIC project, which aims at developing wireless intracerebral probes dedicated to image positron-emitting source activity in the brain of awake and freely moving rats. Former experiments with the PIXSIC positron probe based on a passive sensor have validated the proof of concept, but have also shown limitations with regards to the probe robustness and to its transparency to annihilation photons.

The IMIC circuit features a matrix of  $16 \times 128$  active pixels of  $30 \times 50 \ \mu\text{m}^2$  size and targets to overcome the PIXSIC probe drawbacks by exploiting a thin sensitive layer of 18  $\mu\text{m}$ , still featuring an overall thickness close to 300  $\mu\text{m}$ . Additionally, by using a low power (55 nW/pixel) in-pixel front-end architecture providing binary output, IMIC solves the challenge of implanting an active sensor in tissues where overheating is forbidden.

The needle-shaped sensor 610  $\mu$ m × 12000  $\mu$ m was fabricated and tested in laboratory. The whole sensor dissipates 160  $\mu$ W and its imaging capabilities were asserted with various sources : <sup>55</sup>Fe, <sup>90</sup>Sr and <sup>18</sup>F. These tests also demonstrated robust count-rate measurement with IMIC in the range 10–1000 counts/matrix/s. Finally, a dedicated setup qualitatively confirmed excellent in-sensitivity to 511 keV  $\gamma$ -rays.

In this paper, we present the sensor requirements and its detailed design. We also discuss the first characterisation results and the outlook for the integration of IMIC into an implantable probe.

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Keywords: Semiconductor Detectors, CMOS Pixel Sensors, Beta Probe, Brain Imaging, Neuroimaging

#### 1. Introduction

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Neuroimaging results of anaesthetised or highly restrained 2 animals suffers from undesired effects that are limiting  $_{\scriptscriptstyle 22}$ their validity [1, 2]. In contrast, real time monitoring of  $_{23}$ 4 molecular processes in brains of fully awake animals has a  $_{24}$ 5 strong potential to improve the robustness of observations  $_{25}$ that couple neuroimaging and behavioural studies [3, 4]. 26 Approaches based on PET imaging [5, 6, 7] have only  $_{27}$ 8 partly achieved this goal due to a remaining mechanical 28 9 support, or an overall weight, or a limited space that does  $_{20}$ 10 not guarantee the animal freedom of movement for beha- $_{30}$ 11 vioral studies. 12 31

Another approach consists in performing invasive measurements using an autonomous positron sensitive probe implanted in specific brain regions by stereotaxic surgery. An advanced solution using a needle-shaped passive silicon sensor was developed with PIXSIC [8, 9, 10, 11]. Figure 1 An advanced solution using a needle-shaped passive silicon An advanced solution using a needle-shaped passive silicon and a sensor was developed with PIXSIC [8, 9, 10, 11]. Figure 1 advanced solution using a needle-shaped passive silicon advanced solution using a needle-shaped passive silicon An advanced solution using a needle-shaped passive silicon advanced solution using a ne First the length of the metallic lines that connect pixel sensors to their pre-amplification stages on the PICPUS readout chip induces EM pickup, which reduces signal over noise ratio and hence the overall probe sensitivity.

The second weak point of PIXSIC pertains to the thickness of the passive sensor, which is constrained by antagonist requirements. On the one hand, mitigating the sensitivity to  $\gamma$ -rays generated by electron-positron annihilations, whose positions are not necessarily near the probe, requires a thickness as small as possible. On the other hand, a relatively thick sensor is beneficial to mechanical stiffness. A compromise of 200 µm thickness was chosen for PIXSIC, though

illustrates the full PIXSIC concept. The miniaturisation of the sensor signal treatment within the PICPUS ASIC [9] allows wireless data transmission through a relative light system carried on a backpack. PIXSIC actually releases physical hindrances to the rodent movements. While invasiveness of this technique is a major point of concern, no impact from the small-size probe direct implantation was observed. However, the PIXSIC setup suffers from two main limitations.

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FIGURE 1: Schematic of the PIXSIC setup. The needle probe implanted in the brain is composed of 10 passive pixels, each directly wire-bonded to one channel of the readout chip : PICPUS. This chip embeds amplifiers, filter, digitisation, and memories, and is enclosed in dental cement above the skull. A micro-controller steers and gathers data from the readout chip. It is located on a backpack carried by the rodent during experiments. It also embeds active and passive circuits for the polarisation of the sensor and readout chip, and a battery for full autonomy.

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it was not really satisfactory for both the constraints. 41 42

The usage of monolithic active pixels sensors (MAPS) 78 43 was proposed [12] to improve the implanted probe concept. 79 44 MAPS can embed an amplification and digitisation stage 45 in each pixel, basically replacing the PICPUS chip and 46 suppressing a priori sensitivity to electromagnetic envi-47 ronment. Since the amplification allows a small primary 81 48 signal, the sensitive thickness can be thinner than the ove-  $_{82}$ 49 rall sensor thickness. Hence a depth of typically a few tens  $_{83}$ 50 of micrometers generate enough signal for efficient detec-  $_{84}$ 51 tion and does not need to extent to the full sensor thick- $_{85}$ 52 ness also limiting its sensitivity to  $\gamma$ -rays. The latter can  $_{86}$ 53 be set optimally with respect to invasiveness and stiffness.  $_{\rm sz}$ 54 In addition, the probe integration is simplified due to the  $_{88}$ 55 absence of an additional circuit on the rodent head, as de-56 picted in figure 2. 57 90

However, the usage of an implanted active sensor can  $_{q_2}$ 59 only be effective under the condition that the local tempe- $_{_{93}}$ 60 rature is marginally affected by the probe operation (typi- $_{94}$ 61 cally  $\Delta T < 1^{\circ}$ C). Consequently, a key design feature is the <sub>95</sub> 62 power dissipated by the sensor. 63

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97 Following the ideas developed in [12] along with the  $_{\circ 8}$ 65 requirement to minimize the power consumption, a first  $_{aa}$ 66 MAPS based on the CMOS pixel sensors technology  $has_{100}$ 67 been developed. The chip is named IMIC (Imageur Moléculaire 68 Intra-Cérébral) and is part of the MAPSSIC (MAPS  $\mathit{Sonde}_{\scriptscriptstyle 102}$ 69 Intra-Crânienne) project to develop and exploit the  ${\rm first}_{\scriptscriptstyle 103}$ 70 71 MAPS-based intracerebral positron probe. 104

105 This paper first describes in section 2 the requirements  $_{106}$ 73 for the active sensor to be implanted in the brain. The<sub>107</sub> 74 design and operation features of IMIC, optimized for  $\mathrm{low}_{\scriptscriptstyle 108}$ 75

count rate and power dissipation, are detailed in section 3. Section 4 provides first characterization results, focussing on functional validation and sensitivity with laboratory sources.

#### 2. Sensor Requirements

Neuroimaging exploits a variety of radioactive isotopes, typically <sup>18</sup>F, <sup>15</sup>O and <sup>11</sup>O, which emit  $\beta$ + at energies from 0 to 1.7 MeV. The probe ability to measure activities depends obviously on what energy range the sensor is sensitive to. Consequently an important figure of merit is the detection threshold, which should be as low as possible (this requirement is an outcome of the study in [12]) and depends on the signal-over-noise ratio of the sensing element. The threshold value will be discussed in the next section 3 in view of the sensitive thickness and noise level of the designed sensors.

As previously mentionned, the usage of MAPS for this kind of probe intends to both enhance the overall robustness and simplify the system beyond the implanted part of the probe. Thus, we have included into the requirements the digital conversion of the signal inside the pixels hence facilitating further data transmission. It is also expected that the readout architecture of the sensor optimises data throughput to the micro-controller embedded on the animal backpack with respect to the measured activity.

Regarding the sensors dimensions, we follow the ones of the PIXSIC sensor (15 mm long, 500  $\mu$ m wide and 500  $\mu$ m thick), compatible with the requirements on invasiveness [9].

Imaging the source and evaluating its activity require some level of segmentation, especially along the (vertical) implantation direction. The size of the pixels of the PIXSIC



FIGURE 2: Schematic of the MAPSSIC setup. The monolithic active pixel sensor IMIC is a needle shaped probe composed of  $16 \times 128$  pixels of  $30 \times 50 \text{ µm}^2$ , which provide binary information whether a pixel has been hit or not. IMIC is wire bonded to a PCB acting as an interface to the connector and is enclosed in cement on the head of the rodent. The chip is steered and read out only through digital signals by using a micro-controller located on the backpack together with a battery, which is embedding wireless transmission with the data acquisition PC.

<sup>109</sup> probe were 200  $\times$  500 µm<sup>2</sup>, with no segmentations but<sup>139</sup> along its length. Such a segmentation proved to be granu-<sup>140</sup> lar enough to map subcortical brain activity [9] and there-<sup>141</sup> fore should be matched quite easily by MAPS, which can<sup>142</sup> feature much higher granularity. power of 1 mW per sensor is taken as an indicative limit. The actual guideline chosen for the sensor design follows the requirement to achieve the lowest possible power dissipation.

114 According to first positron count rate observations with  $^{^{143}}\!$ 115 the PIXSIC probe [9], an average positron flux of  $about_{144}$ 116 100 hits/sensor/s is expected over the required dimensions. This level of positron flux corresponds to at least three<sup>145</sup> 117 118 orders of magnitude less than standard achievements of <sup>146</sup> 119 147 CMOS pixel sensors used for charge particle detection. 120 148 However, a stringent constraint arises from operation in 121 149 a living animal brain. 122 150

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As a matter of fact, active architectures involve  $\operatorname{power}^{^{151}}$ 124 consumption and thus heat dissipation, which could locally<sup>152</sup> 125 overheat brain tissues in contact with the sensor. To avoid<sup>153</sup> 126 behavioural impact or worst, tissue damage, from excess<sup>154</sup> 127 heat dissipation, finite element computations were perfor-128 med in order to set a limit on the dissipated power. The si-129 mulated geometry includes a volume of silicon representing<sup>157</sup> 130 the implanted part of the probe  $(10 \text{ mm} \times 500 \text{ } \mu\text{m} \times 600 \text{ } \mu\text{m})^{158}$ 131 in a spheric medium representing the brain at a constant 132 temperature of 37 °C. The highest temperature gradient  $^{160}$ 133 at the direct probe contact stays below the required 1  $^{\circ}\mathrm{C}^{^{161}}$ 134 162 for a dissipated power below 4 mW. Considering the pos-135 sible implantation of multiple probes composed of 2 to  $4^{163}$ 136 sensors, we have set a limit of 1 mW dissipated per sensor.<sup>164</sup> 137 Due to the simplistic nature of the above calculation, the<sup>165</sup> 138

#### 3. IMIC sensor design

### 3.1. Technology choice and overall geometry

We have chosen to design the sensor in the same imager CMOS process with 180 nm feature size as used for various charged particle detection applications [13]. The sensitive volume consist of a 18 µm thick low-doped epitaxial layer, which has demonstrated 100 % detection efficiency for minimum ionising particles for various pixel sizes below 50 µm [13]. Such realization guarantee *a priori* excellent detection performances for  $\beta$ + within the energy range of interest.

In addition, the thin detection layer presents two key advantages. First, it mitigates the sensitivity to the annihilation photons as recommended in [12]. Secondly, it allows thinning the sensor down to thicknesses from 100 to 300  $\mu$ m, thus achieving an optimum between invasiveness and mechanical stiffness.

Regarding segmentation, we have opted for a rectangular shape pixel, within the dimensions known to yield excellent charge collection : 30 µm wide and 50 µm along the vertical dimension of the probe. A matrix of 16 (horizontal) × 128 (vertical) pixels then covers a sensitive area of  $480 \times 6400 \ \mu\text{m}^2$ , meeting the objective to create a



FIGURE 3: Top : layout of the 12000 µm long and 610 µm wide IMIC sensor. Bottom left : magnified layout of a single pixel underlying the location of its main functional components. Bottom right : Pixel circuit schematic derived from the ALPIDE sensor [14].

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needle-shape probe. 167

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The overall sensor dimensions reach 610 µm width and<sub>201</sub> 169 12 mm length. The vertical extension beyond the active<sub>202</sub> 170 matrix serves to implement the readout and control micro-203 171 circuits as well as the connection pads. The sensor is thin-204 172 ned by the foundry to 280 µm and hence presents three<sub>205</sub> 173 different layers along its thickness : first 10 µm of elec-206 174 tronic process insensitive to particles, secondly 18 µm of<sub>207</sub> 175 sensitive epitaxy and lastly roughly 250 µm of silicon sub-208 176 strate that provides mechanical support. 209 177 210

The produced sensor layout is shown on the top of fi-211 179 gure 3. 180 212

#### 3.2. Pixel design 181

To satisfy simultaneously the main requirements on the<sup>215</sup> 182 pixel functionalities and performances (digital output, low 183 data-throughput, low detection threshold and low power),<sup>216</sup> 184 a modified version of the ALPIDE (ALice PIxel DEtec-217 185 tor) [14] circuit developed for the ALICE experiment is<sub>218</sub> 186 implemented. The initial compact pixel architecture am-219 187 plifies, shapes (with clipping to prevent saturation), and<sub>220</sub> 188 discriminates the signal collected by the diode using only<sub>221</sub> 189 12 transistors, as depicted on the bottom of figure 3). 222 190 Pixels integrate any incoming charge between two conse-223 191 cutive readouts. Since the binary output of the front-end<sub>224</sub> 192 indicating the presence of a hit is only available for a few<sub>225</sub> 193 tens of µs, a 1-bit memory was added per pixel to keep this226 194 information until the pixel readout time. The memory is<sub>227</sub> 195 reset after the pixel is read out, so that if a second particle 196 hit the same pixel before readout, the 1-bit memory can-228 197 not account for it. The matrix readout strategy is detailed 198

in the next sub-section.

Simulations of the pixel architecture using Cadence Virtuoso Analog Design Environment have evaluated the equivalent noise charge (ENC) at the collection node below 10 electrons, hence allowing to set discriminators for the digitisation at an equivalent level of  $\approx 100 \text{ e}^-$ . The corresponding energy threshold for the detection equals  $\approx 360$  eV, well below the maximum probability value of energy loss in 18  $\mu$ m of silicon by  $\beta$ + with energies within 0 to 2 MeV.

Finally, these simulations predict that the overall frontend dissipates about 55 nW per pixel (or 113  $\mu$ W for the whole matrix), well below the upper limit. The schematic of the pixel and its layout are shown in figure 3.

#### 3.3. Readout strategy

IMIC readout mechanism is kept simple for the sake of space and power dissipation, which does not depend on the hit rate, and relies on the rolling-shutter principle. A token is propagated through the matrix in a shift register at a fixed clock rate to select one line. The 16-bits word formed by all the pixel memories in the line is serialised towards the output of the sensor at a higher clock rate (at least 16 times), which defines  $f_{clock}$ . This operation is repeated 128 times to gather data from the entire pixel array, thus the time  $t_{readout}$  needed for this full readout cycle amounts to

$$t_{readout} = \frac{2048}{f_{clock}} \quad [s]. \tag{1}$$



FIGURE 4: Readout strategy for IMIC. When a pixel detects a hit, the event is memorised until rolling-shutter readout. A frame is composed of an adjustable waiting time and the readout time. During the readout, the data of each pixel is sent outside the sensor, bit after bit. The data of the memory is reset after readout to allow the detection and memorisation of a new hit.

Pixels integrate particle hits during the time  $t_{wait}$  between two readout cycles, plus the readout time itself:

$$t_{integration} = t_{readout} + t_{wait} \quad [s]. \tag{2}$$

For each readout cycle, data are provided to the micro-232 controller as a vector of 2048 bits to be sent further away 233 through a wireless protocol. In view of limiting the data<sup>273</sup> 234 rate defined by equation 3 and hence the overall power 235 consumption of the MAPSSIC setup, it is better to maxi-236 mise the waiting time  $t_{wait}$  during two readout cycles. Ho-237 wever, avoiding pile-up of hits during the same integration 238 period requires to minimise  $t_{wait}$ . 239

$$Data \ rate = \frac{2048}{t_{integration}} \quad [bits/s] \tag{3}$$

The foreseen average hit-rate of about 100 hits/matrix/s in our experiments, leads to an optimisation for the integration time of around 50 millisecond. In such conditions the data rate stays at a few tens of kilo-bits per second while keeping the pile-up probability at the per mil level. Figure 4 depicts the principle of operation of the IMIC readout.

#### 249 3.4. A fully programmable needle shaped sensor

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In order to minimize the complexity of the external cir-250 cuitry, IMIC limits the number of required connections by 251 using a command decoder controlled through Serial Per-252 ipheral Interface (SPI) protocol. This controller also allows 253 the parametrisation of on-chip digital-to-analog converters 254 (DACs) used for the polarisation of the front-end. The 255 steering signals (and the data) are sent independently bet-275 256 ween the micro-controller located on the rodent backpack<sup>276</sup> 257

and the sensor. The micro-controller connects to the external world through a wireless protocol. The overall system description will be the subject of a subsequent publication.

The digital block driving IMIC is located on the sensor edge near the few connection pads located outside of the brain, opposite to the pixel matrix deeply implanted in the tissues. This block adds a power consumption of approximately 2  $\mu$ W. The simulated total IMIC power consumption amounts to approximately 115  $\mu$ W. This dissipation does not depend on the activity measured but is entirely driven by the number of pixels populating the matrix.

The main characteristics of the IMIC sensor are summarised in table 1.

TABLE 1: Summary of the simulated characteristics of the IMIC sen-

or.	
Feature	Value
Sensor size $[\mu m^3]$	$610 \times 12000 \times 280$
Pixel pitch [µm <sup>2</sup> ]	$30 \times 50$
Sensitive depth [µm]	18
Det. threshold [eV deposited]	$\approx 400$
Pixel power consumption [nW/pixel]	$\approx 55$
Matrix size	$16 \times 128$
Sensitive area [µm <sup>2</sup> ]	$480 \times 6400$
Total power consumption $[\mu W]$	$\approx 115$

#### 4. IMIC characterisation

#### 4.1. First images



FIGURE 5: Images acquired with the IMIC sensor. Single frame aquired with a  ${}^{55}$ Fe soft-X-ray source (5a), a  ${}^{90}$ Sr  $\beta$  source (5b), and a  ${}^{18}$ F positron source (5c). A single annihilation photon from the  ${}^{18}$ F positron source hit (5d). Image composed of 1000 accumulated frames of a metallic spacer acquired with  ${}^{55}$ Fe (5e).

A few IMIC sensors were bonded to a printed circuit board (PCB) in order to facilitate the first tests. After adjustment of the various polarisations through the on-chip

DACs driven by the SPI protocol, the bonded chips were<sup>333</sup>
confirmed to be fully functional. All results reported below<sup>334</sup>
were obtained with these sensors at room temperature and<sup>335</sup>
under relative obscurity.

The measured dissipated power by the whole sensor is 160  $\mu$ W, close to the value expected from simulations in table 1 considering the low absolute value for a complete chip.

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Three types of sources have been used to characterise 288 the IMIC sensor. Single frames from each source type are 289 displayed in figures 5a-5d. Soft X-rays (5.9 and 6.4 keV 290 rays) from the <sup>55</sup>Fe source (figure 5a) generate small clus-291 ters of mainly two pixels, while the  $\beta$  particles, either from 292  $\beta$ - (<sup>90</sup>Sr : figure 5b) or from  $\beta$ + (<sup>18</sup>F : figure 5c) yield 293 clusters about ten times larger in average. This difference 294 comes from the relatively long ionising path of  $\beta$ -rays in 295 silicon. In contrast, X-rays undergo point-like conversion 296 through photoelectric effect. 297

Images from 511 keV  $\gamma$ -rays were obtained by placing the <sup>18</sup>F source far away (see sub-section 4.3) from the sensor. The cluster size observed for these  $\gamma$ -rays is similar to the one for positrons (see figure 5d). This indicates that the cluster size information cannot be used to identify (and suppress) the unwanted  $\gamma$ -rays hits on an individual basis information.

338 To demonstrate that the sensor internal readout per-307 forms as expected, an image of a metallic washer was ac-308 quired using the  ${}^{55}$ Fe source (as shown in figure 5e). The 309 ring is approximately 500  $\mu$ m thick with an external dia-310 meter of 5 mm and an opening diameter of 2 mm. The  $^{342}$ 311 ring structure can easily be recognized on the image as it  $^{343}_{344}$ 312 completely stops the soft X-rays. 313 345

#### 314 4.2. Validation of long integration times

As explained in section 3, the modest awaited activities<sup>347</sup> 315 of the radio tracers allow running the IMIC sensor with  $\mathrm{a}^{\scriptscriptstyle 348}$ 316 long integration time in the order of 100 ms, which in  $\mathrm{turn}^{^{349}}$ 317 reduces the data flow and simplifies the whole system. This<sup>350</sup> 318 readout strategy is made possible by the in-pixel memory<sup>351</sup> 319 and was tested by measuring a range of activities at inte-<sup>352</sup> 320 353 gration times varying from 20 ms to 1 s. 321 For this test, the IMIC sensor was indirectly illuminated<sup>354</sup> 322 with a  ${}^{90}$ Sr  $\beta$  source through an aluminium shield. The<sup>355</sup> 323 activity was modulated using three different  $\operatorname{aluminium}^{356}$ 324 thicknesses : 1, 3 and 7 mm. Additional measurements<sup>357</sup> 325 were performed without the source in order to monitor<sup>358</sup> 326 dark count rate. For each combinations of shield and in-327 tegration time, 1000 frames were acquired with the same<sup>359</sup> 328 detection threshold. Results are presented in figure 6. The<sup>360</sup>

detection threshold. Results are presented in figure 6. The<sup>360</sup> mean entries per integration time were converted into a<sup>361</sup> mean activity measured over the whole matrix : <sup>362</sup>

$$\overline{\mathcal{A}(t_{int})} = \frac{\overline{entries}}{t_{int}} \quad [hits/matrix/s]. \tag{4}_{364}$$

The saturation limit corresponds to the situation where every pixel over the matrix has been fired. This limit depends on the integration time through the simple relation :

$$\mathcal{A}_{sat}(t_{int}) = \frac{2048}{t_{int}} \quad \text{[hits/matrix/s]}.$$
 (5)



FIGURE 6: Mean activity seen by the sensor for integration times between 20 ms and 1 s of the IMIC sensor dark counts, and with various activities of  $^{90}\text{Sr}$   $\beta\text{-rays}$  modulated by using various aluminum shielding thicknesses.

The analysis of the dark count map revealed that a few pixels (denominated noisy pixels in the following) dominate the production of fake hits, which explains the dark count rate evolution with the integration time. As there are one to two noisy pixels systematically firing during one integration period, it follows that the dark count rate simply decreases as  $1/t_{integration}$  along with equation 4. The low level for the dark count rate means that the IMIC sensor can measure robustly activities down to approximately 10 hits/matrix /s.

With the  ${}^{90}$ Sr source attenuated by only 1 mm of aluminum, the mean activity follows the saturation limit curve. At t<sub>int</sub> = 10 ms, the mean activity seen by the sensor is approximately 60000 hits /matrix/s.

Using thicker shielding, the mean activity on the sensor remains below the saturation limit and is constant for every integration time. Thus, in the ballpark of expected average activities (100 hits/matrix/s), integration time up to 1 s can be chosen without information loss and with a safety margin against activity fluctuation of one order of magnitude. This read-out condition allows low data rates, and limits the power consumption of the overall system.

#### 4.3. Preliminary sensitivity study

Measurements using <sup>18</sup>F radioisotope, produced at the CYRCé cyclotron at IPHC, in a non-sealed aqueous solution were performed in view of studying the differential sensitivity to prompt  $\beta$ + and annihilation photons.

A first 3D printed tank contained a diluted solution of  $^{18}$ F

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FIGURE 7: Experimental setup dedicated to measurements with <sup>18</sup>F. The IMIC sensor wire-bonded to its PCB is closely oriented towards a first tank containing an aqueous solution of <sup>18</sup>F. A vial of the same radiotracers with an higher activity was put on demand 15 cm away from the sensor in order to mimic the presence of a parasitic  $\gamma$ -ray source.

presenting an activity of about 1 MBq. The tank was de-376
signed to precisely position the sensor bonded to its PCB377
5 mm away from the solution. Since the source was dissol-378
ved in saline solution, the sensor was protected by plastic379
wrap. 380

Another vial filled with the same isotope but featuring  $a_{381}$ much higher activity of 100 MBq was disposed on demand $a_{382}$ 15 cm away from the sensor. This second tank served as  $a_{383}$ parasitic source of 511 keV  $\gamma$ -rays, since the air distance $a_{384}$ and the glass of the vial (the opening needed to fill the $a_{385}$ vial is not oriented towards the sensor), fully attenuates $a_{386}$  positrons. This setup presented in figure 7 intends to reproduce roughly the head–bladder relative positions in a rat.

Data were taken with the IMIC sensor in three configurations, for 5 minutes each. The observed counts per time and average count rate are plotted in figure 8a, where the average count rates are corrected for the time decay of <sup>18</sup>F. Count distributions observed with both tanks present are labeled <sup>18</sup>F+ $\gamma$ , the distributions obtained with only the tank close to the sensor is tagged <sup>18</sup>F, and finally the label



FIGURE 8: Left : Evolution with time of the pixel count per 100 ms frame duration for the three source configurations described in the text. Right : Average pixel count per frame over five minutes data acquisition for the same configurations.

 $_{387}$  is  $\gamma$  when only the distant tank was used.

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The counting rates for the three source configurations<sub>441</sub> 389 are displayed in figure 8b. Within uncertainties, the ave-442 390 rage counting rate with both sources  $({}^{18}\text{F}+\gamma)$  or only with 443 391 the closer tank (<sup>18</sup>F) are equivalent, they amount respec-444 392 tively to 56.0  $\pm$  0.7 and 57.9  $\pm$  0.8 pixels/frame. This 393 observation is coherent with the very small counting rate 394  $(0.13 \pm 0.04 \text{ pixels/frame})$  measured when only the far-395 away source  $(\gamma)$  is used. 396 446 The experiment illustrates qualitatively the very small re-447 397

<sup>397</sup> The experiment inustrates quantatively the very small let-<sup>398</sup> lative sensitivity of the thin IMIC sensor to  $\gamma$ -rays with <sup>449</sup> respect to  $\beta$ +.

#### 400 5. Conclusion

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Following [12], we have designed a monolithic active<sup>455</sup> 401 pixel sensor to equip the next generation of intracerebral<sup>456</sup> 402 positron probes for deep brain imaging adapted to  $\operatorname{fully}_{_{458}}^{_{457}}$ 403 conscious and freely moving rodents. The IMIC chip fea-459 404 tures 2048 active pixels over a sensitive area of  $480 \times 6400$  mm<sup>2</sup>. 405 The in-pixel amplification and 1-hit binary memory allows<sup>46</sup> 406 both a low detection threshold (400 eV for the energy de- $\frac{1}{463}$ 407 posited) and fully digital matrix readout. This front-end<sub>464</sub> 408 architecture benefits from an extremely low power, addres-465 409 sing the main challenge of heat dissipation within the im-  $^{\rm 466}$ 410 plantation tissues. Finally, being fabricated on a 18  $\mu m_{468}^{VV}$ 411 thick sensitive layer, the sensor is mostly transparent to<sub>469</sub> 412  $\gamma$ -ravs. 470 413

The first IMIC sample characterisations confirmed the  $^{\scriptscriptstyle 471}$ 414 472 successful operation of the sensor as a  $\beta$ + activity loca-473 415 tor. The measured power dissipation per sensor reached<sub>474</sub> 416 160  $\mu$ W. This performance sets the current state-of-the-<sup>475</sup> 417 art for a probe built on the CMOS pixel technology, and  $_{--}^{^{476}}$ 418 should allow the implantation of a few probes in a  $\text{single}_{478}$ 419 animal with acceptable overheating of the local brain tis-479 420 sues. Foreseen *in-vivo* experiments will be conducted to<sup>480</sup> 421 investigate the full compliance of the IMIC sensor with  $^{*\circ 1}_{432}$ 422 the application requirements. 423 483

A second generation sensor can benefit from a fully depleted thin sensitive layer, which would allow to increase the pixel size, hence decrease the total number of pixels and leading to a reduction of the total dissipated power. depleted thin sensitive layer, which would allow to increase the pixel size, hence decrease the total number of pixels and leading to a reduction of the total dissipated power. depleted this sensitive layer, which would allow to increase and leading to a reduction of the total dissipated power. depleted this sensitive layer total dissipated power.

429 Currently, pairs of IMIC sensors are assembled back-to-430 back to form the implantable part of the final MAPSSIC 431 probe with a total volume of  $560 \times 610 \times 12000 \text{ }\mu\text{m}^3$ . The 432 rest of the construction process for the whole probe and 433 its tests in realistic environment will be the subject of a 434 forthcoming publication. 435

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