

Functional Imaging and Instrumentation Group





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"Advances in Position Sensitive Photodetectors for PET applications"

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Outline

PET History

- PET Physics and Technology
- PSDs in PET: PMT
- PSDs in PET: Solid State
- Advanced PET Detectors: DOI and TOF
- SiPMs for PET→ The Ultimate Dream??
- Conclusions



1952 - The beginnings

PET History

- PET Physics and technology
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First Clinical

Positron Imaging Device

1952 - This instrument followed the general concepts of the instrument build in 1950 but included many refinements. It produced both a coincidence scan as well as an unbalance scan. The unbalance of the two detectors was used to create an unbalance image using two symbols to record any unbalance in the single channel rates of the two detectors.





First clinical positron imaging device. **Dr. Brownell** (left) and **Dr. Aronow** are shown with the scanner (1953).

Coincidence and unbalance scans of patient with recurring brain tumor. Coincidence scan (a) of a patient showing recurrence of tumor under previous operation site, and unbalance scan (b) showing asymmetry to the left. (Reproduced from Brownell and Sweet 1953).



1957 - The Anger Camera

PET History

PET Physics and technology

Principle:

- PSDs in PET: PMT
- PSDs in PET: solid state
- Advanced PET detectors: DOI and TOF
- SiPMs for PET
- Conclusions

many photomultiplier tubes "see" the same large scintillation crystal; an electronic circuit decodes the coordinates of each event



Hal Anger (Berkeley) Developer of the scintillation camera



	FIIG			PET spatial resolution / 1
•	PET History	FWHI	M =	$1.2 \sqrt{(d/2)^2 + b^2 + (0.0022D)^2 + r^2 + p^2}$
	PET Physics and technology	(mm)		NonPositronParallaxCrystal sizeCodingcollinearityrangeerror
•	PSDs in PET: PMT	1.2	·	Degradation due to reconstruction algorithm
•	PSDs in PET: solid state	d	:	Crystal pitch
	Advanced PET	b	:	Coding error
	detectors: DOI and TOF	D	:	Detector separation
	SiPMs for PET	r	:	effective source size (including positron range)
•	Conclusions	p	:	Parallax error
		* Dere sens Anal	enzo itivity ysis ii	& Moses, "Critical instrumentation issues for resolution <2mm, high brain PET", in <i>Quantification of Brain Function, Tracer Kinetics & Image</i> Brain PET, ed. Uemura et al, Elsevier, 1993, pp. 25-40.



From man to mice

PET History

- PET Physics and technology
- PSDs in PET: PMT
- PSDs in PET: solid state
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- Conclusions



microPET







*Images courtesy of Simon Cherry, UCLA







FIIG	1986 - The block detector
 PET History PET Physics and technology PSDs in PET: PMT 	In a block detector, a 2D array of crystals are attached to 4 PMTs. Usually the array will be cut from a single crystal and the cuts filled with light- reflecting material. When a photon is incident on one of the crystals, the resultant light is shared by all 4 PMTs. Information on the position of the detecting crystal may be obtained from the PMT outputs by calculating the following ratios and comparing them to pre-set values:
 PSDs in PET: solid state Advanced PET detectors: DOI and TOF 	$R_{x} = \frac{A + B}{A + B + C + D}$ $R_{y} = \frac{A + C}{A + B + C + D}$ V $Crystal cuts form light guides V PMT C$ V
SiPMs for PETConclusions	where A, B, C and D are the fractional amounts of light detected by each PMT

In 1986 the introduction of the block detector by Mike Casey and Ronald Nutt, changed the world of nuclear imaging. *Almost all dedicated tomographs built since1986 have used some forms of the block detector.*







From the block detector to PSPMT's

- PET History
- PET Physics and technology
- PSDs in PET: PMT
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"Block detector"





Large "b"Limitations on minimum "d"

- "1st generation" PSPMT
- Hamamatsu PS-PMT R2486. •50 mm Ø active area • 16 x + 16 y anodes



- Small crystals can be used (down to d = 1mm)

Used in the YAP-(S)PET (Univ of Ferrara Italy,1996)

Flood field irradiation (511 keV) of a matrix of scintillator YAP:Ce, read by a Hamamatsu R2486 (resistive readout)

FIIG	PSPMT's + matrix of scintillators
	1958 IEEE TRANSACTIONS ON NUCLEAR SCIENCE, VOL. 43, NO. 3, JUNE 1996
 PET Physics and technology 	Use of a YAP:Ce Matrix Coupled to a Position-Sensitive Photomultiplier for High Resolution Positron Emission Tomography
PSDs in PET: PMT	A. Del Guerra ^{1,2} , F. de Notaristefani ³ , G. Di Domenico ^{2,4} , M. Giganti ⁵ , R. Pani ⁶ , A. Piffanelli ⁵ , A. Turra ¹ , G. Zavattini ^{1,2}
PSDs in PET: solid state	 ² INFN, Sezione di Ferrara, via Scienze, I-44100 Ferrara, Italy ³ INFN, Sezione di Roma I and Dipartimento di Fisica, Università di Roma III, Italy ⁴ Dottorato di Medicina Nucleare, Catedra di Medicina Nucleare, Facoltà di Medicina e Chirurgia, Corso, Giovecca 203, I-44100
Advanced PET detectors: DOI and TOF	 ⁵ Cattedra di Medicina Nucleare, Facoltà di Medicina e Chirurgia, Corso Giovecca 203, I-44100 Ferrara, Italy ⁶ Dipartimento di Medicina Sperimentale, Università "La Sapienza" Roma, Italy
SiPMs for PET	
Conclusions	



R2486 based small animal scanner YAP-(S)PET II

PET History

- PET Physics and technology
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Hamamatsu R2486 (3" Ø)

YAP:Ce matrix $(4 \times 4 \text{ cm}^2)$ 400 finger crystals 2.0 mm × 2.0 mm × 25 mm





Mouse heart metabolism with ¹⁸F-FDG (2008)

Due to the YAP:Ce crystal and the planar geometry the scanner can perform SPECT too, by adding a parallel hole collimator in front of each matrix

First publication: A.Del Guerra, G.Di Domenico, M.Scandola, G.Zavattini, "High spatial resolution small animal YAP-PET", Nucl. Instr. Methods A, 1998, <u>A409</u>, 537-541. 15



"2nd generation" PSPMT

PET History

- PET Physics and technology
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The PS-PMT R8520-C12 by Hamamatsu.

- Overall size: 25.7 mm × 25.7 mm
- Active area: 22 mm × 22 mm
- 6 x + 6 y anodes

Advantages

- Higher packing fraction with respect to round tubes (up to 73%)
- Good spatial resolution
- Good uniformity
- Easy to use with a resistive chain (4 channels)

Drawbacks

- Small active area for each tube
- Dead area between adjacent PMTs



Flood field irradiation (122 keV) of two matrices of scintillators (CsI:TI, left and YAP:Ce, top) read by a Hamamatsu R8520-C12 (single tube, resistive readout)

Used for small animal imaging!





LSO:Ce – hystorical results

Table 1 Properties of LSO, BGO and NaI(TI)

PET History

- PET Physics and technology
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	LSO	BGO	NaI(TI)
Relat. light intensity	75	15	100
Peak wavelenght	420 nm	480 nm	410 nm
Decay constant (ns)	12 (30%); 42 (70%)	300	230
Density (g/cc)	7.4	7.13	3.67
Effective atomic No.	66	75	51
Index of refraction	1.82	2.15	1.85
Hygroscopic?	no	no	yes
Rugged?	yes	yes	no



This crystal is coupled to a PMT via its 2x2 mm side.

Timing

The coincidence time spread function for LSO and BGO resulted in a FWHM of 0.95 ns for LSO and 2.2 ns for BGO using wo Hamamatsu R647-04 PMTs

FIIG	PSPMT's + fibers + L	SO: MicroPET
	IEEE TRANSACTIONS ON NUCLEAR SCIENCE, VOL. 44, NO. 3, JUNE 1997	1161
PET History	MicroPET: A High Resolution PET Scanner	for Imaging Small Animals
PET Physics and technology	S.R. Cherry ¹ , Member, IEEE, Y. Shao ¹ , R.W. Silverman ¹ , S. Sicgel ¹ , Member, IEEE, A. Chatziioannou ¹ , Member, J.C. Moyers ² , Member, IEEE, D. Newport ² , A. Boutefnouc	Senior Member, IEEE, K. Meadors ¹ , IEEE, J.W. Young ² , W.F. Jones ² , het ¹ , T.H. Farquhar ¹ , Member, IEEE,
PSDs in PET: PMT	M. Andreaco ² , M.J. Paulus ² , <i>Member, IEEE</i> , D.M R. Nutt ² , <i>Fellow</i> , <i>IEEE</i> and M.E	I. Binkley ² , <i>Member</i> , <i>IEEE</i> , 2. Phelps ¹
PSDs in PET: solid state	¹ Crump Institute for Biological Imaging, Dept. of Molecular and Medi ² CTI PET Systems Inc., Knoxvil	cal Pharmacology, UCLA, Los Angeles, CA le, TN.
Advanced PET detectors: DOI and TOF	PS-PMT Hamamatsu R7600-C8	LSO BGO
SiPMs for PET		U Time 2. Desites biogenerate for LSO mission DET design (a)
Conclusions	8 × 8 square fibres bundle (2 mm) 8×8 LSO matrix	and a conventional BGO block detector (right).
		Rat heart 18F-FDG





1952

- PET History
- PET Physics and technology
- PSDs in PET: PMT
- **PSDs in PET:** solid state
- Advanced PET detectors: DOI and TOF
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Scintillators Photodetectors Crystal size Nb of detectors Detector rings **Ring Diameter Animal Port** Field-of-view Nb of slices

Example of application

Initial Results from the Sherbrooke

Avalanche Photodiode Positron Tomograph

R. Lecomte, J. Cadorette, S. Rodrigue, D. Lapointe, D. Rouleau, M. Bentourkia, R. Yao and P. Msaki

Department of Nuclear Medicine and Radiobiology,

Avalanche Photodiode $3 \times 5 \times 20 \text{ mm}^3$ 512 / 32 cassettes (2×8 array) 2 (1 ring of modules) 310 mm 135 mm $118 \text{ mm} \emptyset \times 10.5 \text{ mm}$ 3 (2 directs, 1 cross) 20-40 ns (~25 ns)

BGO

IEEE TRANSACTIONS ON NUCLEAR SCIENCE, VOL. 43, NO. 3, JUNE 1996

Whole-body PET Scan ¹⁸F⁻ + ¹⁸FDG, 250 g rat (Sherbrooke APD PET Scanner)







IEEE TRANSACTIONS ON NUCLEAR SCIENCE, VOL. 46, NO. 3, JUNE 1999 **A** 3D HIDAC-PET Camera with Sub-millimetre Resolution for Imaging Small Animals A.P. Jeavons, R.A. Chandler, C.A.R. Dettmar



SiliPET (proof of principle)

Mesurements with a \approx 1 mm diameter ²²Na source. The MEGA detector was devided into two stacks 2 cm apart made by of 5 and 6 prototype layers. Coincidence events between top and bottom stacks were taken and logical XOR for layers in each stack was implemented. Due to the 1 micosecond time window (MEGA electronics) a low activity source was used to keep random coincidences low.



Reconstructed with a focal plane ray tracing method

0.0 cm

1.0 cm

MEGA prototype tracker: 11 doublesided Si 19 X 19 cm² detector layers. Si wafer parameters: thickness – 0.5mm; strip pitch – 470 microns.



Profile through the sources of the composite image. FWHM = 0.75 mm. Source diameter = 1 mm. Same results are obtained by simulating a uniform 1 mm diameter sphere.



SiliPET (design)

The <u>SiliPET</u> small animal PET scanner is composed by 4 detector stacks each composed of 40 planar, 1 mm thick, **double sided silicon strip detectors** with 128 strips on each side to permit the measurement of the two coordinates on the plane of the detector.

The third coordinate, the depth of interaction, is given by the identification of the detector plane in which the interaction takes place with a precision determined by its thickness.

All planes in a stack are in *exclusive OR* imposing therefore a single interaction.

- □ <u>Stack layer</u> → DOI mesurement
- $\Box \underline{Compton interaction} \rightarrow \text{accurate position}$
- \Box <u>No parallax</u> \rightarrow compact and high sensitivity
- □ <u>No energy mesurement</u> → no ADCs

Di Domenico G, Zavattini G, Cesca N, Auricchio N, Andritschke R, Schopper F and Kanbach G, 2007, SiliPET: an ultra-high resolution design of a small animal PET scanner based on stacks of double-sided silicon strip detector *Nucl. Instrum. Methods Phys. Res.* A **571** 22–5







	FIIG	TOF PET
	PET History	R. Allemand, C. Gresset, and J. Vacher, "Potential advantages of a Cesium Flouride scintillator for time-of-flight positron camera," J. Nucl. Med., vol. 21, pp. 153-155, 1980.
•	PET Physics and technology	Utilizing Photon Time-of-Flight Information
•	PSDs in PET: PMT	Abstract-The physical characteristics and some imaging capabilities of Super PETT I, a positron emission tomograph utilizing time-of-flight
	PSDs in PET: solid state	(TOF) in its image reconstruction process were assessed experimentally by means of measurements carried out in phantoms and clinical imaging studies. The performance characteristics assessed included sensitivity, spatial resolution, image improvements resulting from time-of-flight information utilization, system dead time, and linearity. The clinical examples included imaging of the brain, the heart, the liver, and a
	Advanced PET detectors: DOI and TOF	demonstration of Super PETT Ps capability of achieving cardiac gating. is not entirely settled, there seems to be agreement that a gain
	SiPMs for PET	of several fold (2.5-4) can be obtained with TOF precision measurements of about 400-500 ps, which is practically achievable with modern technology, for objects in size range
	Conclusions	of 30-35 cm, in their transversal dimensions. Fig. 1. Photograph of Super PETT L. CONCLUSIONS AND DISCUSSION
		The following, albeit preliminary, conclusions can be drawn. As predicted, the incorporation of photon time-of-flight
		information into the PET reconstruction process does sub- stantially increase the signal-to-noise ratio in the reconstructed

image for a timing resolution of about 500 ps. The measured









	FIIG	Results: intrinsic timing
		• $\Lambda = 800 \text{ nm}$ • $\lambda = 400 \text{ nm}$
	PET History	Intrinsic timing measured at s.p.e level: 60 ps (σ) for blue light at 4V overvoltage.
•	PET Physics and technology	150 (not subtracted)
•	PSDs in PET: PMT	SiPM illuminated with a pulsed laser with ¹⁰⁰ 60 fs pulse width and 12.34 ns period, with less than 100 fs jitter.
	PSDs in PET:	Two wavelengths measured: $0 \frac{1}{2} \frac{3}{3} \frac{4}{4} \frac{5}{0} \frac{6}{0} \frac{7}{0}$
	solid state	$\lambda = 400 \pm 7 \text{ nm and } \lambda = 800 \pm 15 \text{ nm}.$
•	Advanced PET detectors: DOI and TOF	Time difference between contiguous pulses is determined. $\lambda = 400 \text{ nm}$
		The timing decreases with the number of $_{50}$ at 4 V overvoltage [fit as $1/\sqrt{(N_{c})}$]
	SIPMs for PET	photoelectrons as
	Conclusions	1/√(Npe) → 20 ps at 15 photoelectrons. $_{30}$
		20
		10
		[G. Collazuol et al., VCI 2007, published in NIM A.] $0 \begin{bmatrix} 1 & 1 & 1 \\ 0 & 2 & 4 & 6 & 8 & 10 & 12 & 14 & 16 \\ N_{pe} & N_{pe} \end{bmatrix}$

Results: coincidence timing (TOF)

PET History

- PET Physics and technology
- PSDs in PET: PMT
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SiPMs for PET

Conclusions

 $\sigma \sim \frac{\sqrt{Q} \ \tau}{< N >}$

Coincidence measurement with two

LSO crystals (1x1x10 mm³) coupled

to two SiPMs {Theory: Post and Schiff.

Phys. Rev. 80 (1950)1113.}



Where:

<N> = average number of photons: ~ 100 photons at the photopeak Q = Trigger level: ~1 photoelectron.

 τ = Decay time of the scintillator

For two scintillators in coincidence expected : => $\sqrt{2\sigma}$ ~ 630 ps . <u>Measured => ~ 600 ps sigma.</u>

Measurements in agreement with what we expect!!

[G.Llosa, et al., Conf records of the IEEE NSS-MIC 2007, Honolulu, USA]

Results: energy resolution ($\Delta E/E$)

Setup:

- PET History
- PET Physics and technology
- PSDs in PET: PMT
- PSDs in PET: solid state
- Advanced PET detectors: DOI and TOF

SiPMs for PET

Conclusions

- 2 LSO [1mm x 1mm x 10mm] crystals coupled to 2 SiPMs

- Home made amplifier board.
- Time coincidence of signals.
- VME QDC for DAQ.
- ²²Na source.

Energy resolution in coincidence: 20% FWHM. (best result: 17.5 %)



[G.Llosa et al, Conference Records IEEE NSS-MIC 2006, M06-88]





Position determination

Hitmap for different source position with crystal array







PET/MRI with APDs

¹⁸F-fluorodeoxyglucose - Human

PET History

- PET Physics and technology
- PSDs in PET: PMT
- PSDs in PET: solid state
- Advanced PET detectors: DOI and TOF

SiPMs for PET

Conclusions



SIEMENS



19 x 19 mm crystal block (a): 12 x 12 individual $1.5 \times 1.5 \times 4.5$ mm crystals coupled via a 3 mm thick light guide to a monolithic 3 x 3 APD array (b) (Hamamatsu, Japan)



¹¹C-methylphenidate - Mouse

¹⁸F-fluorodeoxyglucose - Mouse





Results: tests of SIPM in MR system (MRI)

in collaboration with the Wolfson Brain Imaging Center, Cambridge, UK

- PET History
- PET Physics and technology
- PSDs in PET: PMT
- PSDs in PET: solid state
- Advanced PET detectors: DOI and TOF

SiPMs for PET

Conclusions

S.p.e and ²²Na energy spectra acquired with gradients off (black line) and on (red line). No real difference is appreciated in the data.

Differences in photopeak position is due to temp changes in the magnet (apparent change in gain due to changes in breakdown voltage).

Pickup in baseline when switching on/off





100 120 140 160 180

[R.C.Hawkes, et al. to be presented at IEEE, NSS-MIC 2007, Honolulu, USA]

0 20 40 60 80



FIIG	Conclusion #2: photodetectors for small animal PET instrumentation
 PET History PET Physics and technology PSDs in PET: PMT 	Small animal PET instrumentation differs from clinical scanners for the use of <u>higher intrinsic resolution photodetectors</u> , mainly PSPMTs. The <u>one-to-one coupling and DOI</u> measurement are already employed in some small animal scanners
 PSDs in PET: solid state Advanced PET detectors: DOI and TOF 	Solid state detectors could be a valid alternative to the scintillator approach. The <u>multimodality</u> is much required for small animal imaging.
 SiPMs for PET Conclusions 	The <u>SiPM</u> has all of the characteristics: speed, QE, granularity, flexibility, robustness for a successful implementation in small animal instrumentation.



PET Physics and technology

PSDs in PET:

PSDs in PET: solid state

PMT

Conclusion #3

PET History SiPMs and SiPM matrices:

- Are well understood (available, reproducible, robust,...)
- Perform to SPECs (Fast, High PDE, MRI compatible,...)
- Geometry and performance can be tailored to the application!
- Are well suited for PET and also for SPECT applications (both pixellated and slab of scintillator)

Advanced PET detectors: DOI and TOF
BUT..... There is no free lunch!! They NEED:
— Dedicated ASIC (under develor

SiPMs for PET

Conclusions

Temperature control (sensor and feedback on the ASIC)
 Distributed readout (easily 10-50k channel)

Dedicated ASIC (under development)

- Low cost in the future [commercial cost now: ~100\$ per mm²]



Conclusions #4

PET History

- PET Physics and technology
- PSDs in PET: PMT
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- Advanced PET detectors: DOI and TOF
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- Conclusions

Quoted From: "Recent developments in PET detector technology" Tom K Lewellen, Phys. Med. Biol. 53 (2008) R287–R317

"......Further, my crystal ball predicts that the winner for the ultimate PET detector design is likely to be a mosaic detector made up of elements consisting of slabs of crystal (perhaps 50 x 50 mm²) viewed by arrays of SiPMs and supported by statistical-based estimation algorithms that locate events in the crystal slabs in three dimensions....."

→The design as in ref: "A detector head design for small-animal PET with silicon photomultipliers (SiPM)",S. Moehrs, A.Del Guerra, et al. Phys. Med. Biol. 51(2006) 1113–27





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