

## EVALUATION AND DEVELOPMENT OF QUALITY CONTROL PROCEDURES FOR TECHNETIUM-99M LABELLED MACRO AGGREGATED ALBUMIN AT A TERTIARY HOSPITAL IN GAUTENG

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The radiopharmaceutical, technetium-99m labelled macro aggregated albumin (99mTc-MAA) is an injectable diagnostic agent used in Nuclear Medicine for lung perfusion imaging in patients with suspected pulmonary embolism.

Ventilation and lung perfusion imaging scans (V/Q scans) are done routinely in the Nuclear Medicine Department at Dr. George Mukhari Academic Hospital (DGMAH) to assess patients with suspected pulmonary embolism.

Little documented information exists about the quality control procedures of 99mTc-MAA for lung perfusion imaging at the tertiary hospital.

The need for evaluation and development of the quality control procedures is based on establishment of a new Single Photon Emission Computed Tomography (SPECT) unit at the tertiary hospital in 2019.

In practice, when validated quality control procedures such as those in the pharmacopoeias and in the accepted guidelines are impractical to perform in a hospital radiopharmacy due to various reasons such as technical complexity of the procedures, expensive analytical components, time consumption, toxicity of solvents, alternative quality control procedures can be developed that are faster, simplified and cost effective. For this reason, 99mTc-MAA is one of the radiopharmaceuticals currently being used at the hospital hence there is existing need for evaluation and development of quality control procedures for the radiopharmaceutical.

The main objective of this study is to validate current quality control processes for 99mTc-MAA at the tertiary hospital's Nuclear Medicine Department, to identify any missing or undetermined quality control processes and to develop these quality control procedures according to the relevant guidelines.

The research methodology is quantitative, analytical and experimental. The methods involve experimental tests on the radiopharmaceutical and these are; radionuclidic purity, radiochemical purity, particle size, particle number and pH test.

The outcome of this study is to develop a quality control tool for 99mTc-MAA where necessary and to validate the existing quality control procedures on the radiopharmaceutical.

The implication of this study to other researchers is the provision of a validation method for quality control procedures of radiopharmaceuticals. Newly set-up hospitals can use the validation methods in this study, as well as alternative quality control procedures that this study seeks to develop.

Keywords: Technetium-99m labelled macroaggregated albumin, validation, quality control tool, development, quality control procedures.

# VALIDATION OF A GAS CHROMATOGRAPHY METHOD FOR THE DETERMINATION OF RESIDUAL SOLVENTS IN PET RADIOPHARMACEUTICALS

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## Background

PET radionuclides have a short half-life and, unlike conventional pharmaceuticals, must usually be prepared, tested, and administered to patients within a very short time. The current European Pharmacopeia (EP) method for determining residual solvents using gas chromatography flame ionization detection (GC-FID) is tedious and time-consuming. Apart from a long analysis time of  $\pm 60$  minutes, some peaks were not well defined on chromatograms and peak tailing occurred. The validation of non-pharmacopeial analytical procedures is necessary to demonstrate that they are fit for the intended purpose.

## Aim

The purpose of this research was to validate a much faster in-house developed method, using the International Conference on Harmonisation for Technical Requirements for Registration of Pharmaceuticals for Human Use (ICH) guidance document.

## Methods

Standard solutions were prepared in-house using the highest grade ethanol, methanol, and acetonitrile.

The method was then validated in accordance with the ICH guidelines. Parameters that were tested included specificity, precision, accuracy, linearity, range, limit of detection, limit of quantification, and robustness. Linearity was established over a concentration range of 50% to 150% of the ICH limits for all the solvents.

## Results

The in-house developed method had a total analysis time of 8 minutes which is significantly shorter than the method described in the EP. The method was also found to be specific, accurate, precise, and linear. The method was able to detect and quantify all the residual solvents present in [<sup>18</sup>F]FDG and [<sup>18</sup>F]FMISO samples. The limit of detection (LOD) and limit of quantification (LOQ) were found to be 75ppm and 750ppm for methanol; 250ppm and 1000ppm for ethanol; 20.5ppm and 102.5ppm for acetonitrile respectively. The method was also found to be precise, with intra-day and inter-day relative standard deviations (RSDs) of less than 3%. Accuracy was determined by recovery of the samples, ranging from 99.5% to 99.7%.

## Conclusion

An in-house method for determining residual solvents in PET radiopharmaceuticals was successfully tested and validated using ICH guidelines. The results showed that this method decreased the analysis time significantly and can be used on a routine basis.

## THE INFLUENCE OF LOW- DOSE CT ATTENUATION CORRECTION ON ARTEFACTS OF MYOCARDIAL SPECT IMAGES FOR NUCLEAR MEDICINE STUDIES

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**Introduction:** Artefacts in myocardial perfusion SPECT imaging cause poor image quality, which may present as false-positive results that may lead to unnecessary, expensive and invasive procedures. Breast and diaphragm attenuation may cause anterior and/or inferior soft tissue artefacts and may interfere with the visualisation of the perfusion defects of the myocardium. The myocardial perfusion studies performed with Tc-99m sestamibi, comprised of stress/ rest images with patients positioned in a supine or prone position.

**Aim:** The aim of the study was to determine whether hybrid imaging, SPECT with computed tomography (CT), will improve the image quality by reducing the soft tissue artefacts with the application of attenuation correction maps in the stress and rest Tc-99m sestamibi myocardial SPECT perfusion studies.

**Materials and Methods:** A retrospective study was performed on 100 image data sets of patients with suspected ischaemic heart disease referred to the Nuclear Medicine Department. The no attenuation corrected (NAC) and the attenuation corrected (AC) stress and rest images were the dependent variables of the study. The independent variables were the relevant factors such as gender, age, weight, height and position of the patients during imaging were also reviewed for their influence on the attenuation correction of the images. The results of the patients were quantitatively reviewed and the images were given a score of 0-4. An overall score of the results of the improvement (yes) versus no improvement (no) of attenuation correction application (AC-A) was also given. The critical assessment areas were the inferior and anterior wall defects and analysis was performed by the two Nuclear Medicine Physicians. A Generalized Linear Model was applied to determine relationships between the dependent and independent variables.

**Results/ Discussion:** The effect of the independent variable on the dependent variable with a p-value of >0.05, indicated non-significance. The results demonstrated that with anterior stress and rest SPECT images, the most artefacts were created with AC-A. Improvement for the stress inferior images was 61%. AC-A to the stress and rest attenuation corrected (AC) and (NAC) images scored an improvement of 49%. Although no improvement of 51% for the overall outcome of stress/rest studies, it included 18 stress studies had improvement and nine rest studies improved (scores 2-3) in the inferior wall of the myocardium.

**Conclusion:** The AC-A scores of the images for AC versus NAC with MPI showed a significant influence on the outcome of the study on the inferior wall of the myocardium.

## Radiolabelling of Chelators DOTA, EDTA and DTPA with Gallium-68

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### Introduction:

The use of [68Ga]Ga-EDTA, [68Ga]Ga-DTPA and [68Ga]Ga-DOTA in cisternography has been described. If such products are to be used in a clinical setting, reliable production and quality control methods are essential.

### Aim:

The aim of this study was to validate iTLC and HPLC methods to determine radiochemical purity of selected chelators and to translate manual methods into a suitable automated radiolabelling method.

### Methods:

Manual radiolabelling of chelators DOTA, EDTA and DTPA with gallium-68 was performed as previously described. Gallium-68 was obtained from an iThemba LABS germanium-68/gallium-68 generator less than 1 year old using pre-purification or fractionated elution. Labelling was performed at pH 4.5; incubated at 95°C for 10 min. Solid-phase extraction was used to remove colloids from the final product whereafter a quality control sample was obtained for iTLC and HPLC analysis. A two-strip SG-iTLC method was validated with a range of concentrations using acetonitrile and TFA as mobile phases. A radio-HPLC method was developed (Phenomenex Jupiter 4µm Proteo 90A Column, 250 x 4.6 mm) using gradient elution conditions with acetonitrile and deionised water and flow rate 1 ml/minute. The optimal labelling parameters determined by manual synthesis were translated to an automated synthesis method using the Scintomics GRP 3V module.

### Results:

Fractional elution or pre-purification followed by 1000 µl of sodium chloride elution resulted in the highest concentration of Ga-68 for radiolabelling of chelators. iTLC analysis showed free Ga-68 at solvent front ( $R_f = 0.4 - 0.6$ ) in TFA whereas free gallium-68 and colloid remained at the origin ( $R_f = 0$ ) in acetonitrile. A solvent mixture of 50 % acetonitrile and 4 % TFA resulted in optimal peak resolution for [68Ga]Ga -EDTA and [68Ga]Ga -DOTA whereas for [68Ga]Ga -DTPA the solvent mixture consisted of 20 % acetonitrile and 4 % TFA. HPLC analysis revealed a retention time of 4.34-4.58 min for both free gallium-68 and radiolabelled 68Ga-chelators with no other radiochemical impurities present. Automated radiolabelling of [68Ga]Ga -DOTA showed a radiochemical purity > 95% on iTLC analysis.

### Conclusion:

The optimized radio-iTLC method showed separation of colloids, free gallium-68 and radiolabelled chelators. HPLC analysis may require a different selection of stationary and mobile phases for separation of radiochemical species. Formulation requirements relevant to intrathecal administration of radiopharmaceuticals will be addressed in future studies.

## Dacryoscintigraphy in a traumatic lacrimal duct injury.

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### Introduction

Epiphora may be secondary to problems associated with secretion or excretion of the lacrimal system. It is important to know which of these problems are associated with a patient's presentation, as the management of both problems are different. Ultrasonography can be used to produce images of the lacrimal sack and the flow of tears; however, the ultrasonic probe exerts pressure on the medial canthal area (potentially affecting pressure gradients created under physiological conditions). Dynamic MRI studies of lacrimal drainage are possible but challenging to perform.

Dacryoscintigraphy, a non-invasive, functional examination of the lacrimal system uses technetium-99m pertechnetate to evaluate tear flow through the lacrimal drainage system. Nasolacrimal duct obstruction is a common problem, especially in children. Trauma to the canaliculus most often leads to epiphora. Repair of the canaliculus lessens the risk of the patient developing epiphora.

Dacryoscintigraphy offers the referral doctor the opportunity of knowing if the epiphora is due to an obstruction, and it can further determine at what level the obstruction has taken place. It has been extensively used to facilitate the definite diagnosis of obstructions and stenosis of the lacrimal drainage system. Dacryoscintigraphy is easy to perform and is non-invasive, and advantage It has over contrast dacryocystography, thus avoiding general anaesthesia in children.

### Methods

A dacryoscintigraphy was performed on a 7-year-old patient who sustained a dog bite injury to the face two years prior, and now presenting with right eye epiphora. The patient received 1 drop of 100uCi Tc-99m Pertechnetate to the lateral canthus of each eye. Dynamic images followed by serial static images were acquired.

### Result

In the left lacrimal system, immediate visualisation of the lacrimal sac, with spontaneous drainage into the lacrimal duct was observed. There was progressive accumulation of the radiotracer in the lacrimal sac of the injured right eye, up to the end of the study, without any notable drainage into the nasolacrimal duct noted. The study concluded that there was evidence of obstruction at the level of the left lacrimal sac.

### Conclusion

Dacryoscintigraphy mimics "physiological" lacrimal outflow and can be performed under normal pressure gradients, it is considered more suitable for the study of functional epiphora. In this clinical case it confirmed that the epiphora was due to an obstruction and this was able to guide the next appropriate management in this patient.

## Can FDG PET/CT be used to optimize the treatment of patients with pulmonary tuberculosis?

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Can FDG PET/CT be used to optimize the treatment of patients with pulmonary tuberculosis?

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### Background:

The treatment of pulmonary tuberculosis is long due to the slow-growing nature of *Mycobacterium tuberculosis* (MTb). At the end of a standard course of anti-tuberculous treatment (ATT), some patients still harbour slow-growing MTb (sgMTb) in their lesions, which are responsible for relapse. These sgMTb are non-culturable and are difficult to detect. We investigate the utility of end-of-treatment [18F]F-FDG PET/CT to differentiate patients with sterilizing cure versus those with sgMTb who are at risk for relapse.

**Methods:** We prospectively recruited patients who had a standard course of ATT. All patients underwent [18F]F-FDG PET/CT within two weeks of completing ATT. We determined the presence of residual metabolic activity (RMA), which is FDG uptake in the lungs above mediastinal background activity on the [18F]F-FDG PET/CT. Patients were classified as having RMA or complete metabolic response (CMR) to ATT. All patients were subsequently followed-up after completing [18F]F-FDG PET/CT imaging for relapse. Confirmatory bacteriologic testing was performed in patients with clinical suspicion of relapse. In those who relapsed, a repeat FDG PET/CT imaging was subsequently performed.

**Results:** We studied 75 patients including 50 HIV-infected individuals with a mean age of 36.09 (SD 10.49). The median CD4 count among HIV-infected patients was 255 cells/mm<sup>3</sup> (IQR: 147– 448). HIV viral load was 12497 copies/mL (IQR:158-38841). HIV-infected patients had lower hemoglobin levels (13.07±1.78 g/dL versus 14.24±2.07, p=0.021) and higher C-reactive protein levels (5.70 versus 1.20,

p=0.001) compared with HIV-uninfected patients. All other baseline clinical and demographic characteristics were not significantly different between the groups. Forty-one patients had RMA and its incidence was not significantly different between HIV-infected and uninfected patients (p=0.101). Thirty-four patients demonstrated complete metabolic response (CMR) to ATT. No tuberculosis relapse was demonstrated in those with CMR. In the RMA group, 3 patients (7.3%) relapsed on follow-up. Relapse was confirmed bacteriologically in all the 3 patients. Repeat [18F]F-FDG PET/CT imaging at the time of clinical relapse demonstrated the persistence of RMA in the same lung regions in all three patients.

Conclusion: FDG PET/CT imaging demonstrates a high prevalence of RMA among patients treated with a standard course of ATT for PTB, with a similar incidence between HIV-infected and uninfected patients. CMR on post-treatment [18F]F-FDG PET/CT is consistent with sterilizing a cure for pulmonary tuberculosis. Patients with RMA on post-treatment [18F]F-FDG PET/CT are at risk for tuberculosis relapse. These results show the potential of [18F]F-FDG PET/CT for use as a non-invasive biomarker for drug development and in investigating shorter ATT regimens.

## RADIOPHARMACY SERVICES IN SOUTH AFRICAN ACADEMIC HOSPITALS AND IMPLICATIONS FOR PLANNING NEW SERVICES.

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**Introduction:** Nuclear medicine (NM) plays an important role in diagnosis and treatment of a wide range of diseases. Optimal radiopharmacy services are key to progress in both patient care and research. Underdeveloped countries are embracing NM as a speciality. To establish new NM and radiopharmacy centres, proper planning is essential due to the high costs of staffing, equipment and radiopharmaceuticals. This study describes the pattern of radiopharmacy services and staffing in NM departments in South African Academic Hospitals (SAAHs); this information can help plan new nuclear and radiopharmacy centres in similar settings.

**Objectives:** The primary objective was to determine the utilization of radiopharmacy services in NM departments in SAAHs and to develop a tool to help plan new radiopharmacy centres in African settings.

**Method:** The study was retrospective (2022), descriptive and quantitative with qualitative aspects. An anonymised questionnaire was sent via Survey Monkey® to the nine SAAHs, to determine hospital and specialist unit sizes, NM patient numbers by scan type, identify facilities (cameras and equipment), staffing levels, commonly used radiopharmaceuticals and to define basic requirements for establishing new facilities.

**Results:** The initial response rate was 67% (6/9 hospitals). Preliminary results: Hospital sizes ranged from 846 to 3200 beds, average NM patients seen was 3,526 for 2022. All hospitals have SPECT-CT cameras; five have PET-CT cameras and one refers PET patients to a private hospital. Only two hospitals have hot cells. All have NM physicians (range 3-13, average 5) and at least one medical physicist (range 1-4) plus nuclear medicine technologists (range 8-15, average 9). Only two hospitals have radiopharmacists (one each) and they also have hot cells. Staff, equipment, patient load and radiopharmaceuticals prepared will be related to patient specialist unit sizes and to catchment population to provide guidelines for start-up services.

**Conclusion:** To promote optimal patient services and support research in SAAHs, norms must be developed that are applicable to Africa. The two hospitals with hot cells have radiopharmacists, indicating these staff are key to progress. Study findings will be compared with IAEA staffing norms to provide recommendations for African settings. Appropriate staff quotas and equipment are required to promote optimal use of radiopharmaceuticals and enable the NM team to operate to their full potential.



## RADIOPHARMACEUTICALS ANALYSIS: BIOMEDICAL CHROMATOGRAPHY APPLICATIONS

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Radiopharmaceutical synthesis methods use relatively large amounts of reagents, necessitating the use of high-resolution and high-capacity chromatographic methods. Furthermore, purification is required prior to the formulation of radiopharmaceuticals for injection to remove various impurities. However, for routine injectable production, radio-thin layer chromatography (i-TLC) is preferred due to short time runs and user friendliness. The problem with i-TLC is that TLC plates are not as efficient as high-performance liquid chromatography (HPLC) columns and are also limited to radioactive detection, so impurities are frequently missed. In this study, the aim was to develop and validate shorter liquid chromatography methods utilizing Gabi, a diode array (UV-VIS), and an evaporative light scattering detector (ELSD) all in conjunction. The analytes of interest in this study were <sup>68</sup>Ga-Fapi, <sup>68</sup>Ga-Dotatate, and <sup>68</sup>Ga-PSMA-617. Degradation of chelators and the introduction of impurities can occur during radiopharmaceutical synthesis. If the sample contains chromophores, the UV-VIS detector can detect them all. The Gabi is capable of detecting all substances containing radionuclides that emit gamma rays. All substances that can diffract light can be detected by the ELSD detector. In this study, it was demonstrated that a simple (5–10 minute) HPLC analytical method with three simultaneous detection systems can be more profitable than the conventional iTLC method for determining the purity of radiopharmaceuticals.

## VISUAL AND QUANTITATIVE COMPARISON OF DIFFERENT IMAGING RECONSTRUCTIONS ON A CLINICAL ACQUISITION

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### ABSTRACT

#### Problem Statement

Filtered backprojection (FBP) is the most basic image reconstruction algorithm used for positron emission tomography (PET). It is also the algorithm of choice to compare acquisition quality between vendors. Advances in system hardware and software now allow for various processing algorithms and applications to be used. These include iterative algorithms (OSEM/MOSEM), point-spread-function (PSF), and time of flight (TOF) applications. Each of these applications can affect quantifiable data obtained from clinical acquisitions. Showcasing the quantitative differences using alternate reconstruction algorithms when viewing 18F-fluorodeoxyglucose (18F-FDG) will benefit troubleshooting in a clinical setting and highlight the importance of consistent image reconstruction.

#### Methodology

Retrospective patient data acquired on a Siemens Biograph Vision 450 PET/CT scanner, with a 64-slice computed tomography (CT), was used. Four (n=4) 18F-FDG scans were reconstructed using five reconstruction techniques (FBP, OSEM with and without TOF, MOSEM with and without TOF), with CT attenuation correction. SUVmean for target organs (brain, heart, liver) was obtained as well as SUVmean for blood pool background (superior vena cava). Target organ to blood pool ratio was obtained (SUVR) and averaged over the four acquisitions for each of the five reconstruction methods.

#### Expected Findings and Conclusion

As expected, differences in PET-CT reconstruction influenced factors such as SUV values (or SUVR) and effects visual changes. Therefore, it would be advised to identify an optimal reconstruction method and apply it to all scans acquired at a given facility as it also influenced the visual assessment of the images by the physicians.

In conclusion, choosing suitable reconstruction parameters is critical for producing high-quality PET-CT images. A comprehensive understanding of the influence of these parameters on image quality can aid in optimizing the imaging process, ultimately improving diagnostic accuracy.

## Developing a radiolabelled tracer for prospective PET imaging of *Plasmodium falciparum* infection

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### Background

*Plasmodium falciparum* (*P. falciparum*) causes the most severe cases of malaria. Severe malaria (SM), an understudied multisystem disease, affects the host's organs and can lead to serious complications, some effecting life-long neurological and cognitive sequela. A major limitation of understanding SM is the inability to visualise the host-parasite mechanisms governing pathogenicity in situ. Nuclear imaging could aid in investigating malaria-related pathologies, however, there are no malaria-specific tracers. Therefore, we aimed to develop zirconium-89 (<sup>89</sup>Zr) and gallium-68 (<sup>68</sup>Ga)-radiolabelled malaria-specific tracers, assess their stability, in vitro characteristics and in vivo biodistribution using non-invasive microPET/CT imaging.

### Methods

A *Plasmodium*-specific antibody (IIIB6), antibody fragment (Pf-Fab), and peptide (P1) were investigated. IIIB6 and Pf-Fab were conjugated to p-isothiocyanatobenzyl-desferrioxamine followed by complexation of <sup>89</sup>Zr. [<sup>89</sup>Zr]Zr-IIIB6 and [<sup>89</sup>Zr]Zr-Pf-Fab were injected into healthy BALB/c mice and assessed by microPET/CT imaging and post-mortem biodistribution. A 1,4,7,10-tetraazacyclododecane-1,4,7,10-tetraacetic acid (DOTA)-Gly-functionalized P1 derivative (DOTA-P1) underwent <sup>68</sup>Ga-radiolabelling. Radiolabelling was optimised by adjusting radiolabelling reaction conditions. Radiolabelling efficiency (LE), radiochemical yield, and stability was assessed with radio-HPLC and iTLC. In vitro studies for [<sup>68</sup>Ga]Ga-DOTA-P1 included plasma stability, red blood cell (RBC) binding, and plasma-protein binding.

### Results

Quantification of the microPET/CT image-guided cardiac region over 24 hours indicated pharmacological half-lives of 7.1-12.1 h and 3.5-6.0 h for [<sup>89</sup>Zr]Zr-IIIB6 and [<sup>89</sup>Zr]Zr-Pf-Fab, respectively. Unfavorably-high concentration of [<sup>89</sup>Zr]Zr-IIIB6 in liver occurred after 2-6 h and moderate spleen>kidneys>heart>stomach>lung>femur uptake. For [<sup>89</sup>Zr]Zr-Pf-Fab, unwanted kidney uptake occurred after 4-6 h, and moderate uptake in liver>lung>stomach>femur. Ex vivo biodistribution studies confirmed high thermodynamic and physiological stability of both tracers and no organ toxicity. DOTA-P1 was successfully radiolabelled with <sup>68</sup>Ga and purified yielding a radiochemical purity exceeding 97%. Efficient preparations (including purification) of [<sup>68</sup>Ga]Ga-DOTA-P1 (LE=77-99%) at a concentration of 22.7 nmol/ml was achieved after 30 min incubation at 95 °C. This optimized radiosynthesis achieved a radiochemically-pure product (>99%) of adequate molar activities (>1.0 GBq/μmol) for preclinical application. [<sup>68</sup>Ga]Ga-DOTA-P1 maintained stability in human plasma up to 2 h (>90%), showed moderate RBC association (±25%) and high serum protein binding (±87%).

### Conclusion

The development of three radiotracers for future malaria-specific imaging is presented. The results support that [<sup>89</sup>Zr]Zr-Pf-Fab has the more preferable pharmacological profile over [<sup>89</sup>Zr]Zr-IIIB6 and is suggested for future investigation in *P. falciparum*-infected animals. Further tests of [<sup>89</sup>Zr]Zr-Pf-Fab should include co-administrations of compounds that may reduce its unwanted kidney retention. The robust radiolabeling protocol along with its high serum stability and low-level RBC binding supports future pre-clinical characterisation of [<sup>68</sup>Ga]Ga-DOTA-P1 in vivo.

## Radiation exposure of Nuclear Medicine radiographers and radiopharmacists during their PET-CT rotation

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**Purpose:** Radiographers and radio-pharmacists working at a PET CT facility are among the professions most exposed to radiation in nuclear medicine. This study assessed the exposure of our staff during a period of 6 consecutive months to evaluate their exposure per day and per examination and to compare it with published data

**Material & Method:** Seven direct reading dosimeters, by different suppliers, were issued to eight radiographers, and three to five radio-pharmacists. Qualified and student professionals were involved. More than one radiographer and one radio-pharmacist were working practically every day. Only vendor supplied <sup>18</sup>F labelled tracers were used.

The following data were entered by each personnel on personalized paper documents: date, person's identity, dosimeter number, start reading, closing reading. After transcription on Excel, calculation of the individual daily dose and sorting by date was performed. Cross-correlation with the PET database allowed the calculation of the dose per PET examination.

**Results:** There were 140 entries for radiographers and 112 entries for radio-pharmacists.

In the case of radiographers, occasionally high exposures resulted in higher average exposure (28,5 uSv/day). When these peaks were excluded, the average exposure per patient was 17,5 uSv/day or 4 uSv/PET exam. Radio-pharmacists had an average daily exposure of 13,7 uSv/day

Several issues were noted. The transcription of paper data into Excel was time-consuming and prone to errors. The collection of data was irregular over the period of investigation. All dosimeters were not necessarily equally calibrated, with results either in milliSievert or in microSievert. Some personnel seemed more prone to higher exposure, although in rather exceptional circumstances.

**Conclusion:** The exposure recorded at our institution remained within the limits of the IRCP and compared favorably with the figures recorded by others.

Among the improvements suggested by this study, are: the direct entry of data in Excel, regular review of the readings, compliance to existing standard operating procedures, calibration of the dosimeters in the same units, supervision of junior staff, and appropriate shielding (portable screen or injection box) and/or an automated injector.

In addition, TOF cameras and data recovery by artificial intelligence are designed to limit radiation exposure to both patients and staff and allow the recording of PET images with much lower activity.

## SYNTHESIS, RADIOLABELING AND PRELIMINARY IN VITRO ASSESSMENT OF NOVEL RADIOPHARMACEUTICALS AS POTENTIAL BACTERIAL-SPECIFIC RADIOTRACERS USING POSITRON EMISSION TOMOGRAPHY IMAGING.

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**Background:** The current PET-radiopharmaceuticals used for diagnosis of bacterial infections rely on secondary host-mediated inflammatory responses, leading to false-positive findings due to misinterpreted, sterile inflammation. To discriminate between infection and inflammation, a more direct targeting mechanism for bacteria is desirable. Therefore, the development of bacteria-selective PET-radiotracers is a valid approach to improve differential diagnosis of bacterial infections and monitoring the performance of treatment interventions using non-invasive nuclear imaging. In this study three candidate molecules with known bacterial-specific interaction were deemed appropriate radiotracer vector scaffolds when functionalized with 1,4,7-triazacyclononane-1-succinic acid-4,7-diacetic acid (NODASA) to subsequently allow for gallium-68 complexation: (1) NODASA-D-lysine (NDL) and two Mycobacterium Tuberculosis (MTB)-specific phage display-generated peptides also known as (2) NODASA-PH1, and (3) NODASA-H8.

**Methods:** Solid phase peptide synthesis (SPPS) was used to synthesize these compounds, including NODASA-functionalized L-lysine (NLL; negative control), followed by purification using reverse-phase preparative HPLC, and characterization using HRMS/LCMS. Compounds were radiolabeled with generator-eluted <sup>68</sup>Ga-activity, the radiosynthesis was optimized by adjusting reaction conditions such as pH, temperature, and NODASA-compound concentration. Radiolabeling efficiency, radiochemical yield and radiochemical purity was assessed with radio-HPLC or TLC. The radio-metal complex stability was assessed using the EDTA trans-chelation challenge. In vitro studies included quantification of plasma stability, blood cell association, and plasma-protein binding assays to characterize the compound stability, radioisotope integrity and estimated bioavailability following intravenous administration into human blood as a potential PET radiotracer.

**Results:** NDL, NLL, H8 and PH1 were successfully synthesized and conjugated to NODASA, purified (>95%), and characterized. Compounds were subsequently radiolabeled with gallium-68, optimal labeling conditions were explored to produce radiochemically pure products (>99%) including suitable radiochemical yields and activity concentrations for future preclinical applications. All radiometal complexations demonstrated good integrity when challenged by 1000-times excess of EDTA (>90%). [<sup>68</sup>Ga]Ga-PH1, [<sup>68</sup>Ga]Ga-NDL and [<sup>68</sup>Ga]Ga-NLL were stable for up to 2 hours (>90%) in human plasma whereas <sup>68</sup>Ga-H8 degraded rapidly (<40% remained intact after 30 min). All compounds showed minimal RBC interaction (<10%) and plasma protein binding ([<sup>68</sup>Ga]Ga-NDL/-NLL <14%; [<sup>68</sup>Ga]Ga-PH1/-H8 <9%), thereby mimicking a favorable expected bioavailability.

**Conclusion:** Following successful radiolabeling and in vitro characterization, [<sup>68</sup>Ga]Ga-NDL and [<sup>68</sup>Ga]Ga-PH1 are deemed promising candidate PET-radiotracers. Upcoming bacterial cell uptake assays and microPET/CT imaging studies in mice will further assess the potential of these compounds as bacterial-specific ([<sup>68</sup>Ga]Ga-NDL), or even MTB-specific ([<sup>68</sup>Ga]Ga-PH1) radiotracers.

## RE-TREATMENT OF BENIGN THYROID DISEASES WITH RADIOACTIVE THERAPY.

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### Background:

Benign thyroid diseases are the most common thyroid conditions seen at our department. These include Graves' disease, diffuse toxic goiter, toxic or nontoxic goiter and solitary hyperfunctioning thyroid nodules. Treatment options are antithyroid treatment, radioactive iodine and total or subtotal thyroidectomy. Radioactive iodine therapy has been used to treat benign thyroid diseases for many years since the 1970s. Low dose of I-131 is used for treatment of benign thyroid diseases.

### Aim:

The aim of this study was to assess the effectiveness of I-131 therapy in benign thyroid diseases in our setting.

### Methods:

A retrospective analysis was carried out for patients with benign thyroid conditions who underwent single or double I-131 therapy for benign thyroid disease at our department between January 2009 and December 2021. The thyroid function status of the patients was also documented prior to discharge.

### Results:

A total of 127 patients were included in the study. The patients included in the study had the following conditions: Graves' disease 81% (n=104), toxic multinodular goitre 10% (n=13), toxic adenoma 6% (n=7), Marine Lenhard syndrome 2% (n=3) and autonomous nodule 1% (n=1). They were treated with an empirical dose of I131, ranging from 10mCi and 40mCi. Seventy-four percent (n=94) of the patients responded to the single RAI therapy (n=33 euthyroid; n=61 hypothyroid). The remaining 26% (n=33) required retreatment as they remained hyperthyroid. Second RAI therapy was administered to 88% (n=29) of patients while 12% (n=4) of the patient defaulted treatment. The majority of retreatment patients had Graves' disease 90% (n=26), 10% (n=3) had toxic multinodular goitre and 3% (n=1) had toxic adenoma. On discharge 66% (n=83) of the patients were hypothyroid and 31% (n=40) were euthyroid. Three percent (n=4) of the patients defaulted treatment before hypothyroid /euthyroid state could be achieved.

### Conclusion:

Graves' disease patients showed majority of treatment failure compared to other benign thyroid conditions. This is contrary to the commonly reported literature which state that multinodular goiter is more resistant than Graves' disease. Hypothyroid state was archived in more than half of patients.

Keywords: RAI therapy, Hypothyroid, Euthyroid and Graves' disease.

## PERSISTENT SESTAMIBI RETENTION ON DUAL PHASE PARATHYROID SCINTIGRAPHY IN HASHIMOTO'S THYROIDITIS - A CASE REPORT

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### Introduction:

Parathyroid imaging plays a critical role in preoperative management of parathyroid lesions. (1) It localizes hyperfunctioning parathyroid tissue, and specifically parathyroid adenomas with a high sensitivity and specificity. (2)

Parathyroid lesion detection with dual-phase Technetium-99m (Tc- 99m) Sestamibi (MIBI) scintigraphy is dependent of the differential washout of the tracer from the thyroid parenchyma. In some cases, such as in autoimmune thyroid disease (AITD) or in the case of type-1 amiodarone induced thyrotoxicosis (AIT-1), scintigraphy may show persistent uptake of the thyroid parenchyma, with no or minimal washout of tracer on delayed imaging, posing a challenge for interpretation. (3)

### Presenting case:

53-year-old female known with Hashimoto's thyroiditis (HT) referred for new onset hyperparathyroidism. Ultrasound had features of HT, bilateral hypodense thyroid nodules and bilateral lower pole parathyroid glands. Dual-phase parathyroid scintigraphy showed diffuse uptake in the thyroid gland parenchyma, with persistent tracer retention on delayed imaging.

### Method:

A combination of dual-tracer – dual-phase imaging with single photon computed tomography/computed tomography (SPECT/CT) with Tc-99m MIBI and subtraction using Tc-99m pertechnetate was performed.

### Outcome:

Dual-phase Parathyroid scintigraphy showed diffuse uptake with persistent tracer retention in the thyroid gland parenchyma on 3-hour imaging. A bulky, elongated left thyroid lobe was identified and no ectopic focus of abnormal uptake were seen. SPECT/CT reconstructed images confirmed the findings.

Thyroid scintigraphy showed an enlarged gland with inhomogeneous tracer uptake. A bulky left inferior pole and an impression of 'cold' nodules in the left superior pole and right inferior pole were appreciated.

Based on the subtraction analysis, a diagnosis of parathyroid lesion in the inferior pole of the left and right lobes were considered and ultrasound correlation ± FNA was recommended for 'cold' nodule correlation.

### Discussion:

Literature has described focal thyroid disease, subacute thyroiditis, AITD such as Graves' disease, hypertrophic form of HT, as well as AIT-1, where MIBI is not equally washed out on delayed imaging. Specifically, AITD is a significant factor for the retention of MIBI. Very few cases exist in the literature of Hashimoto's thyroiditis as a cause of persistent MIBI retention. (3,4)

Theoretically, the above-mentioned factors increase false negative findings, as a result of reduced visibility of small parathyroid adenomas or parathyroid hyperplasia. (3)

### Conclusion:

This case demonstrated that AITD such as HT should be included in the list of differentials for persistent thyroid gland retention of MIBI, resulting in decreased sensitivity for the visualization of hyperfunctioning parathyroid lesions on dual-phase parathyroid scintigraphy.

# LATE PRESENTATION OF CONGENITAL HYPERINSULINAEMIA DIAGNOSED ON A 18-FLUORO-L-3,4-DIHYDROXYPHENYLALANINE POSITRON EMISSIONS TOMOGRAPHY SCAN - FIRST CASE AT UNIVERSITAS ACADEMIC HOSPITAL

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## Introduction

Congenital hyperinsulinism (CHI) is a clinical entity that presents as persistent hypoglycemia in infants and children. Congenital hyperinsulinism classically presents at birth and poses a high risk for hypoglycemic brain injury and neurodevelopmental abnormalities; thus, swift diagnosis and treatment are extremely important. Although the classical presentation is at birth, CHI covers a wide clinical spectrum, often leading to presentation after infancy.

Under normal physiological conditions, insulin secretion is typically suppressed if the blood glucose concentration is lower than 5.0 mmol/l.

In patients with CHI, this normal physiological pathway is disrupted, resulting in inappropriate insulin secretion with a failure to suppress insulin secretion.

Diagnosis typically relies on a combination of clinical evaluation, laboratory investigations, molecular genetic testing, and advanced imaging modalities such as 18-fluoro-L-3,4-dihydroxyphenylalanine positron emission tomography (18-F-DOPA) PET/CT scan.

The role of 18-F-DOPA in CHI mainly centers on confirmation of diagnosis with or without molecular genetic testing and in distinguishing diffuse disease from a localized pancreatic disease. The latter plays an essential role in deciding the treatment path for the patient, as patients with focal disease benefit from surgical management and those with diffuse disease benefit from medical treatment with drugs like diazoxide.

We present our first case of CHI at Universitas academic hospital. An 8-month-old female patient was referred from Bongani regional hospital with a history of persistent hypoglycemia and recurrent diffuse tonic-clonic seizures. This late presentation of CHI posed a diagnostic dilemma because, classically, patients with CHI will present around birth with symptoms. After initial laboratory and conventional radiological investigations were inconclusive, the patient was referred to the Department of Nuclear Medicine for F-18-DOPA PET/CT.

## Method

We retrospectively reviewed the findings of the 18-F-DOPA PET/CT study, and reviewed the images for the intensity and pattern of uptake.

## Results

PET/CT Images of the abdomen and pelvis revealed higher than normal physiological uptake throughout the pancreas, with the head having the highest intensity. No other abnormal findings were noted.

## Conclusion

This case report shows the value of F-18-DOPA PET/CT in the confirmation of CHI. The diffuse nature in this particular case prompted the referral doctors to commence medical treatment and not surgical treatment. The patient was subsequently initiated on a trial of Diazoxide and Hydrochlorothiazide for five days. After showing an initial positive response, the child was discharged.



## GA-68 DOTATATE AVID ECTOPIC ADRENOCORTICOTROPIC HORMONE SECRETING PULMONARY CARCINOID TUMOUR DETECTED ON PET/CT

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Tumours causing ectopic Cushing's syndrome (ECS) are rare and cause excess cortisol as well as adrenocorticotrophic hormone (ACTH) over-production with distinct clinical features. Pulmonary carcinoids tend to be more common than other causes of ectopic ACTH syndrome. Ectopic ACTH tumours are often not visible with conventional imaging, however depending on the histology, Gallium-68-DOTATATE, DOTATOC, and DOTANOC positron emission tomography/computed tomography reportedly exhibits greater sensitivity in identifying an ectopic ACTH tumour source.

Here we present a 39 year male who presented with biochemical abnormalities and new onset diabetes and hypertension; a diagnostic work up for Cushing syndrome was undertaken; and it was found that 24-hour urine free cortisol, midnight cortisol and ATCH levels were elevated. The MRI showed no pituitary lesion and CT was unremarkable. This alluded to an ACTH –dependant aetiology and the patient was referred for a [68Ga] -DOTATATE PET/CT scan to assist in the detection and/or localization of a possible ACTH-dependent primary lesion. The scan showed pathological uptake in a parenchymal nodule in the right lower lobe, thought likely to represent the ACTH-secreting tumour. This was confirmed on histopathology post surgical intervention. Patient had resolution of symptoms and was placed on surveillance. A repeat [68Ga] -DOTATATE PET/CT scan was done 4 months later to assess for residual tumour which revealed residual post-surgical inflammatory changes in the region of the right lobectomy.

Ectopic ACTH secretion from pulmonary carcinoid tumours (typical or atypical) can present as a challenge in diagnosis and management; however [68Ga]-DOTATATE can be useful as an initial diagnostic modality in this rare condition which has a high mortality and morbidity.

## DOES BONE SCINTIGRAPHY CHANGE CLINICAL MANAGEMENT IN PATIENTS WITH LUNG CANCER

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Lung cancer is the third most common cancer in South Africa men, with a lifetime risk of 1 in 98. According to the ESMO cancer guidelines, the standard work up includes computed tomography, Xrays, PET/CT and bone scintigraphy.

Bone scintigraphy is commonly used in oncology for assessment of primary and secondary skeletal malignancies. In cancers that are classically characterised by osteoblastic skeletal metastases, bone scintigraphy is known to have high sensitivity for skeletal metastases. This is in contrast to other malignancies that typically have osteolytic bone metastasis or mixed osteolytic/osteoblastic skeletal lesions such as lung carcinoma. The diagnostic yield for these scans is lower on bone scintigraphy. The aim of this retrospective review is to evaluate the utility of bone scintigraphy in management of patients with lung carcinoma.

### Materials and methods

A review was carried out from patient records at Inkosi Albert Luthuli central Hospital, Nuclear medicine department from the years 2017 – 2022. The study was done as retrospective analysis of patients with biopsy proven lung cancer who were referred for bone scan. Patients all underwent bone scintigraphy in the nuclear medicine department at Inkosi Albert Luthuli Central Hospital. Image acquisition was performed on a Siemens biograph SPECT/CT camera. The CT scan was unenhanced and CT parameters were 120kev 50mAs. The bone scan was acquired 2-4 hours after an injected dose of 20mci of 99mTc-MDP. Regional SPECT/CT of the area of interest was acquired at 3min/bed position.

### Results

The study had 40 participants, 33 males and 7 females, with an age ranging from 45 to 83 years. Histology demonstrated non-small cell carcinoma for all of the participants, with the exception of two patients (who demonstrated adenocarcinoma). Our results demonstrated that bone scintigraphy influenced patient staging in 25% of cases and changed management in 17.5% of the cases. Serum ALP results were available in only 8/40 (20%) patients and was increased in all of these patients. (Detailed statistical analysis to follow).

### Conclusion

In our study population, bone scintigraphy changed staging in 25% of patients with lung cancer and influenced management in 17.5%. Bone scintigraphy may not be the optimal nuclear medicine study and further studies are needed to compare the use of bone scintigraphy to that of 18F-FDG PET/CT in lung cancer patients.

## THE CASE OF MULTIPLE PERIARTICULAR MASSES ON BONE SCAN

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Tumoral calcinosis is a rare disorder that was initially described as an arthropathy of calcium pyrophosphate dihydrate deposition. It tends to occur more predominantly in the periarticular location of the extensor aspects of the large joints. It is a benign familial metabolic dysfunction of phosphate regulation, resulting in high levels of phosphate in the plasma leading to calcific deposits of hydroxyapatite or amorphous calcium phosphate crystals into soft tissues. It can also result from other conditions such as renal failure, dialysis, and hypervitaminosis D.

The index case is a patient operated in 2019 for tumoral calcinosis of the right knee, now referred to our department with painless periarticular swellings of the right femur and right elbow.

Biochemically, the patient was noted to have an elevated phosphate level, with normal calcium and parathyroid hormone (PTH) levels. Technetium-99m Methylene Diphosphate (99mTc MDP) bone scintigraphy was undertaken, confirming the findings of tumoral calcinosis in the suspected sites, as well as identifying other sites of involvement.

The use of bone scintigraphy has been proven to have a good sensitivity, and its ability to scan the whole body in one acquisition gives the modality the added advantage of identifying tumoral calcinosis skip lesions. This was well demonstrated with the above case where the patient was treated with multiple surgeries based on radiological findings which were isolated to the area of complaint. Bone scintigraphy was able to identify multiple skip lesions (inclusive of bone marrow involvement), which resulted in change of the management approach in the patient.

Bone scintigraphy is therefore highly recommended, especially in patient with tumoral calcinosis, as it is able to identify lesions both in the suspected and other (skip lesion) sites.

## EVALUATION OF THE ADDED VALUE OF A DIURETIC RENOGRAM IN THE ASSESMENT OF RENAL FUNCTION PRIOR TO TARGETED RADIONUCLIDE THERAPY

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Introduction and study aim:

Lutetium -177 (Lu-177) is a radionuclide with unique theranostic properties that make it ideal for radionuclide therapy. The use of Lu-177 has shown promising results in the treatment of different types of cancers, including neuroendocrine tumors and prostate cancer.

In order for patients to qualify for Lutetium -177 radionuclide therapy a renogram to assess kidney function and rule out obstruction is required. Other tests including a Ga -68 PSMA is also done.

This study aimed to compare the effectiveness of Ga – 68 PSMA and Dotatate derived split renal function with diuretic renogram based split function in assessing the renal function.

A retrospective analysis (visual and quantitative) was conducted on 21 patients with histologically confirmed prostate ca who are Metastatic castration-resistant prostate cancer (mCRPC) and 10 patients with histologically confirmed well-differentiated NET- or moderately-differentiated NETs, G1 or G2 and not amenable to curative surgery. Baseline renal function, renogram parameters and 68 Ga PSMA and Dotatate kidney findings were assessed. Renogram, Ga-68 PSMA and Ga -68 Dotatate were done within a period of +/-1 week of each other. Diuretic renogram split function was calculated with symbia net software.

To calculate relative Ga-68 PSMA and Dotatate distribution in both kidneys, volumes of interest were drawn and generated using syngo.via software. The volume and the mean standard uptake value (SUVmean) were then used for calculating the split renal function (SRF).

of the 31 patients with a mean age of 61years two were noted to have a difference in split function of more than 10% between the diuretic renogram and Ga-68 PSMA and Dotatate derived split renal function. The balance of the patient's demonstrated congruence between the diuretic renogram and Ga-68 PSMA and Dotatate derived split renal function. Concordance was also noted between renal abnormalities on diuretic renogram and Ga-68 PSMA and Dotatate scans.

The results showed that Ga-68 PSMA and Dotatate derived split function had a high correlation with diuretic renogram based split function in assessing renal function. Moreover these tests also provided additional information regarding the functional status of the kidneys.

Apart from assessing function of the kidneys, anatomical defects can also be visually assessed.

Therefore it is suggested that Ga- 68 PSMA and Dotatate derived split renal function can be used as an alternative to diuretic renogram based split function.

## IRRADIATION OF YTTRIUM MICROSPHERES AT THE IRT-T REACTOR

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National Research Tomsk Polytechnic University (TPU), together with an industrial partner (BEBIG), start produced yttrium microspheres at its research nuclear reactor this fall. This radiopharmaceutical is intended for the treatment of liver cancer in inoperable patients and, unlike analogs, it destroys the tumor in a targeted manner without affecting healthy organs and tissues. Irradiation of yttrium-89 microspheres will be carried out at the IRT-T reactor, power 6 MW, neutron flux  $1.7 \cdot 10^{14}$  neutron/cm<sup>2</sup>. To obtain one dose of the final preparation, a weighed portion of 0.1 g of yttrium microspheres is irradiated. Microspheres are processed using distilled water, alcohol and hydrochloric acid. Yttrium microspheres are injected into the patient's bloodstream, which delivers them directly to the tumor. After delivery, the microspheres block the access of blood with oxygen to the metastases, in parallel acting on them with beta radiation. In the Russian Federation, this method of treatment is not yet massively applied.

The industrial production of yttrium microspheres began in the fall of 2020. There have already been test deliveries to Moscow clinics. The drug is registered and has permits for use in clinical practice. April 9, 2021 at A. Tsyb Medical Radiological Research Center for the first time in Russia, clinical trials of the method of radioembolization of tumors with domestic microspheres produced by the Russian company "Bebig" began. Four operations were performed at once on patients with inoperable forms of liver cancer.

## Physiological Biodistribution of Ga-68 DOTA-NOC in Patients with Neuroendocrine Tumours: Influence of Blood Glucose Level, Somatostatin Analogue Therapy and Imaging Time

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Background

A common feature of neuroendocrine tumours (NETs) is that they express somatostatin receptors (SSRTs). Radiotracers such as Ga-68 DOTA-NOC, Ga-68 DOTA-TATE and Ga-68 DOTA-TOC are SSRT analogues and have proven very useful in imaging patients with NET.

SSRTs are also found in a number of normal tissues. Differentiating pathological uptake from normal uptake in organs can be challenging when interpreting Ga-68 DOTA scans. Detailed reference ranges for the physiological uptake in organs has been published for Ga-68 DOTA-TATE and Ga-68 DOTA-TOC, but data are scarce for Ga-68 DOTA-NOC.

SSRT receptor expression by normal organs demonstrates some intra-individual variation and may be affected by blood glucose levels, use of somatostatin analogue therapy or the delay in imaging time.

The aim of this study is to provide detailed reference ranges for the physiological uptake in organs for Ga-68 DOTA-NOC and compare these with limited prior literature. In addition, we will evaluate the effect of blood glucose, somatostatin analogue therapy and imaging time delay on normal Ga-68 DOTA-NOC uptake.

### Methods

Patients having undergone Ga-68 DOTA-NOC PET/CT scans at the NuMeRI Node for Infection Imaging PET/CT centre between 2019 and 2022 will be retrospectively selected. In patients with more than one Ga-68 DOTA-NOC scan only the first study will be included. The clinical history, blood glucose level, any recent somatostatin analogue therapy and the time delay between injection and imaging will be documented.

Uptake measurements will be captured by drawing 3D volumes of interest (VOI) in a number of organs. The maximum standard uptake value (SUVmax) for each organ will be documented, making sure that the VOI is drawn within the organ boundary and excludes any vascular structures. Organs with known tumour involvement will be excluded.

### Results

We expect to provide detailed reference ranges for physiological organ uptake of Ga-68 DOTA-NOC. It is expected that certain values will be higher than what was previously documented in literature. e.g. for the uncinate process. We also expect that blood glucose levels, somatostatin analogue therapy and delay in imaging time to have a measurable effect on the physiological Ga-68 DOTA-NOC uptake in organs.

### Conclusion

More comprehensive reference ranges for physiological organ uptake of Ga-68 DOTA-NOC and improving our understanding of the effects of blood glucose levels, somatostatin analogue therapy

and delay in imaging time on normal organ uptake, we expect to enhance the interpretation of Ga-68 DOTA-NOC PET/CT studies.

## NEEDLE IN A HAYSTACK- MALIGNANT INSULINOMA

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### Introduction:

An insulinoma is a neuroendocrine tumour localized in the pancreas that produces excessive amounts of insulin. They are usually benign and are a rare occurrence with approximately four cases per million. Malignant insulinomas are remarkably rare accounting for 10% of all cases. Nuclear medicine plays a useful role in the localization of insulinomas using Gallium68 Dotatate Positron Emission Tomography /computerized tomography (PET/CT).

### Case Study:

A 45-year-old female with a history of Retroviral Disease presented with recurrent hypoglycaemia. A CT scan of the abdomen showed liver lesions with no pancreatic lesions or insulinoma. She was then referred for a Gallium68 Dotatate PET/CT in May 2020. The patient underwent a Whipple and hepatectomy. She had a follow-up Gallium68 Dotatate PET/CT in April 2021.

### Results:

The initial Gallium68 Dotatate PET/CT done in May 2020 showed avid disease in the body of the pancreas as well as a large liver lesion. After undergoing surgery, a CT scan done in March 2021 showed mesenteric lymph nodes and multiple liver lesions. This prompted a second Gallium68 Dotatate PET/CT in April 2021 for restaging and recurrence. The scan showed extensive hepatic and skeletal disease as well as possible recurrence in the body and tail of the pancreas.

### Conclusion:

Due to the size of insulinomas as well as the fact that they can occur in any part of the pancreas, they are often hard to detect. Gallium68 Dotatate PET/CT scans are very useful in this regard because of their high affinity to somatostatin receptors. This makes it a viable option for insulinoma localization within the pancreas and identification of other sites of metastases as demonstrated in this case study. Furthermore, it can improve diagnostic accuracy and surgical planning for patients with this rare and elusive tumour.



## DIFFERENCES IN TUMOUR AGGRESSIVENESS BASED ON MOLECULAR SUBTYPE AND RACE MEASURED BY [18F]FDG PET METABOLIC METRICS IN PATIENTS WITH INVASIVE CARCINOMA OF THE BREAST.

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### Abstract

**Aim:** To assess 2-deoxy-2-[18F]Fluoro-D-glucose ([18F]FDG) positron emission tomography with computed tomography (PET/CT) semi-quantitative parameters in locally advanced invasive ductal carcinoma (IDC) of the breast and the differences in these parameters based on molecular subtype and race.

**Methods:** [18F]FDG PET/CT images of women with treatment-naïve locally advanced IDC of the breast were reviewed. Qualitative reading of the scans was done to determine presence of regional and distant metastases. Semi-quantitative reading of the scans was done by generating volume of interest using iso-contouring and a fixed relative threshold of 40% and the maximum standardized uptake value-body weight (SUVmax), metabolic tumour volume (MTV), and total lesion glycolysis (TLG) of the primary breast lesion were obtained. The hormone receptor status, HER2 status and biodata including race were retrieved from their medical records. Statistical analysis for differences in the parameters based on race and molecular subtype was done. We categorized the patients by their self-identified race as Black, Mixed ancestry, or White.

**Results:** A total of 127 staging [18F]FDG PET/CT were included comprising of 81(63.8%) Mixed ancestry, 40(31.5%) Black and 6(4.7%) White patients. Mean age was 51.5 years (SD 12.78) and there was no significant difference in age between the various groups. In the IDC group as a whole, the primary tumour SUVmax, MTV and TLG as well as presence of distant metastases was significantly higher in black patients ( $p= 0.004, 0.04, 0.005, 0.002$  respectively). The primary tumour SUVmax, MTV and TLG were significantly higher in basal subgroup ( $p < 0.001$  in all). The primary tumour SUVmax and presence of distant metastases were significantly higher in Black patients in the luminal subgroup ( $p= 0.012$  and  $0.006$  respectively), the SUVmax and TLG were significantly higher in Black patients in the basal subgroup ( $p= 0.057$  and  $0.007$  respectively).

**Conclusion:** Significant differences in [18F]FDG PET/CT parameters were seen in molecular subtype of IDC corresponding to known aggressiveness. The significantly higher parameters in Black patients in IDC as a whole and in luminal and basal subtypes is suggestive and supportive of a more aggressive disease.

## MILK SCAN - DOES LONGER SCANNING TIME INCREASE THE DETECTION OF REFLUX

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### OBJECTIVE:

The milk scan is a well-established technique for detecting gastro-oesophageal reflux (GER) and pulmonary aspiration. There is no universally accepted protocol for this study. At RXH, the dynamic imaging protocol for milk scans in search for GER has been to image for 30 minutes at 5 seconds/frame. However, internationally, imaging is acquired over 60 minutes at 5 seconds/frame. This bodes the question of whether we may have missed refluxes by not scanning for a further 30 minutes, thereby reducing the sensitivity of our protocol.

### METHOD:

We retrospectively reviewed 200 patients who presented for milk scans over a 13-month period from 1 November 2021 to 30 November 2022, 100 patients imaged with the 60-minute protocol, and 100 patients imaged with the 30-minute protocol. We compared the number of positive scan results in the 60-minute protocol to the positive ones in the 30-minute protocol. Then, we further analysed how many of the positive scans in the 60-minute protocol showed reflux in the last 30 minutes of the scan. Has the 30-minute protocol significantly reduced the sensitivity of the reflux search?

### RESULTS:

In the patients scanned with the 60-minute protocol, 65/100 patients had positive scans in comparison to 70/100 patients imaged with the 30-minute protocol in the 13-month review period. In the patients that demonstrated positive studies in the 60-minute protocol, 7/65 patients demonstrated reflux in the last 30 minutes of the study and 21/65 patients demonstrated reflux in the first 30 minutes and did not demonstrate reflux in the last 30 minutes, 37/65 patients demonstrated reflux throughout the 60-minute study. In this review period of 13 months, there was no increase in the incidence of detecting reflux in our milk scans over the 60-minute period in comparison to the 30-minute period.

### CONCLUSION:

Increasing the milk scan protocol to 60 minutes did not demonstrate an increase in the incidence of detection of reflux in our center. The significance of patients in whom reflux was detected in the last 30 minutes of the 60-minute study needs to be reviewed in further studies.

## Lutetium-177 peptide receptor radionuclide therapy of de-differentiated thyroid cancer

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### INTRODUCTION

Well differentiated thyroid cancer when originates from thyroid follicular epithelium and retains its biological characteristics. This allows for it to be treated with radioactive iodine therapy(RAI). About 5% of patients develop metastatic thyroid disease which refractory to RAI. Lutetium-177 labelled peptide receptor radionuclide therapy (PRRT) was explored as an alternative option.

### Methods

63 year old female with follicular thyroid cancer post thyroidectomy more than 10 years ago, presented with right hip pain due to a surgically irresectable mass in the right hemi-pelvis with lytic destruction of the right hip involving the lesser trochanter, acetabulum, ilium and pubic rami.

A diagnostic iodine-123 (I-123) whole-body scan (WBS) revealed uptake in the thyroid bed (thyroid remnant tissue), focal uptake in the left cervical lymph node and heterogenous uptake in the right hemi-pelvis, most intense over the right hip lesser trochanter. she received 2 cycles of I-131 therapy in 2019 at 150mCi and 2021 at 200mCi. Post 2nd cycle of I-131 therapy, the post therapy no longer showed uptake in the right pelvic mass with exception of a focus in the lesser trochanter. A de-differentiated thyroid carcinoma was suspected.

F18-FDG PET/CT scan revealed a destructive, locally infiltrative lesion in the right hemipelvis with abnormal, intense and homogenous uptake. A Ga-68-DOTATATE PET/CT scan demonstrated somatostatin receptor expression at a Krennig score 4 in the mass, congruent to the FDG uptake, with a mildly avid right inguinal lymph node. Post workup for PRRT, she received a total of 4 cycles of Lu-177-DOTATATE therapy.

### Results:

Post 3 cycles, her ECOG score improved from 3 to 1 and her post therapy scans showed a reduction in the size and uptake of the pelvic mass. The post therapy scan following her 4th cycle revealed stable disease. The Ga-68 DOTATATE was repeated as per protocol, at 6 months after her fourth cycle. This scan revealed progression of the lesion in right hemipelvis with regard to intensity and size.

### Discussion and conclusion

Treatment options for de-differentiated thyroid cancer refractory to I-131 are limited and tend to show a suboptimal response. PRRT with beta-emitting and alpha- emitting radionuclides serves as an alternative option in thyroid cancer lesions expressing somatostatin receptors. The relatively short range of Lu-177 may be a factor which results in a slow response given the size of the lesion of the above patient, and tandem treatment with Y-90 PRRT may need to be considered.

## POTENTIOMETRIC STUDIES OF $^{117}\text{MSn}$ -POLYHEDP AND $^{177}\text{LU}$ -POLYHEDP: IN ATTEMPTS TO DEVELOP EFFECTIVE RADIOPHARMACEUTICALS FOR BONE METASTASES

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Secondary cancer tumour formation, often called metastases, remains one of the great scientific challenges in public health. Therapeutic radiopharmaceuticals for bone pain palliation play an important role in providing quality of life for cancer patients with metastatic bone cancer. They comprise of a bone seeking bisphosphonate ligand and a radionuclide. The structural variations of the bisphosphonate affects to a great extent the effectiveness of the radiopharmaceutical with the greatest shortfall being bioavailability. A polymer ligand poly-HEDP was synthesized from its free acid form in relatively low yields. This study seeks to understand the in vivo chemistry of the  $[^{117}\text{mSn}]\text{Sn(II)}$ ,  $[^{117}\text{mSn}]\text{Sn(IV)}$ , and  $[^{177}\text{Lu}]\text{Lu(III)}$  complexed with poly-HEDP in blood plasma. The formation constants of Sn(II), Sn(IV), and Lu(III) were measured by glass electrode potentiometry at 25 °C and I = 150 mM. This made possible the construction of a blood plasma model of poly-HEDP, determined with the aid of thermodynamic blood plasma modeling simulations. The Sn(IV)-poly-HEDP complex was shown to be unstable, with a 100 % dissociation, while Sn(II)-poly-HEDP showed much improved stability with 10 % of the metal ion remaining bound to the ligand. However, 97% of the Lu(III) remained complexed to the ligand with less competition from the physiological ligands in vivo. The synthesis of poly-HEDP from poly acrylic acid was achieved successfully. A thermodynamic blood plasma simulation revealed that Lu(III) is practically quantitatively complexed to poly-HEDP as compared to Sn(II) and Sn(IV) in vivo.

## FDG-PET/CT imaging of Tuberculosis in children

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**Background:** Tuberculosis (TB) causes significant morbidity and is a leading cause of death globally. Young children are at higher risk for severe forms of TB, including disseminated disease and TB meningitis. Up to a fifth of patients with TB in high-incidence countries are younger than 16 years, with a high mortality in untreated children. The diagnosis of childhood TB is challenging as the disease is often paucibacillary, and sampling of specimens difficult, causing poor sensitivity of available pathogen-based tests. Tuberculosis can mimic numerous other pathologies, complicating differentiation between these, active TB or latent disease. The diagnosis of active TB is crucial to initiate adequate treatment timeously, avoid complications and prevent transmission. Evaluation of CXR has several limitations, therefore cross-sectional imaging, e.g., CT, should be considered in the diagnostic evaluation of a symptomatic child. Positron emission tomography combined with (x-ray) computed tomography (PET/CT) using fluorine-18 fluorodeoxyglucose (FDG) has been shown to detect active TB with accuracy equal or superior to other conventional imaging modalities and its role in complicated TB is expanding. FDG PET/CT can effectively identify sites of intrathoracic and extrathoracic TB, assess disease activity, assist in differentiating between active and latent disease, monitor response to therapy, and identify potential biopsy targets. There is a paucity of literature on PET/CT in childhood TB. The efficacy of FDG PET/CT in the paediatric population will be the focus of this case series.

**Methods:** A retrospective case series of children between the ages of 3 months to 16 years who presented at Tygerberg hospital between January 2000 and August 2020 and who had a FDG PET/CT scan for suspected tuberculosis will be performed. The PET/CT scans will be reviewed by a qualified nuclear physician and compared to the findings of the CT scan (if available) and Chest X-ray (CXR). Clinical data will be collated from patient folders, and bronchoscopy and laboratory databases. The data will be analysed using standard descriptive and non-parametric statistical techniques.

**Results:** FDG PET/CT was used mainly for 4 indications: to assist in the diagnosis of MDR TB, identify suitable biopsy sites, evaluate for ongoing disease in complicated TB, and for disseminated disease. The demographic profile and PET/CT findings (including their impact on management) will be presented.

**Conclusion:** FDG PET /CT is expected to impact on the management of selected paediatric tuberculosis cases by demonstrating extent and activity of disease, identifying extrathoracic sites of disease and guiding biopsies.

## The role of [18F]FDG-PET/CT imaging in selecting candidates for radionuclide therapies targeting PSMA

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### Background

Tumor sites with low PSMA-avidity and high FDG-avidity represent sites of aggressive disease which cannot be effectively targeted by radioligand therapy. The presence of discordant lesions seen in very advanced mCRPC has a poorer prognosis. The VISION and TheraP trials where patients were required to have highly PSMA avid disease with no discordant lesions on [18F]FDG-PET/CT prior to [177Lu]-PSMA therapy showed higher response rates. Ideally, all patients who are potential candidates for PSMA targeted radionuclide therapies should undergo [18F]FDG PET/CT together with [68Ga] PSMA PET/CT to detect sites of low PSMA-avidity and high FDG-avidity which preclude the use of targeted by radioligand therapy, however this is not always possible due to the high cost involved. The aim of this study was to assess the impact of [18F]FDG-PET/CT in selecting patients eligible for PSMA targeted radionuclide therapy.

### Methods

Twenty (20) patients with metastatic castration resistant prostate referred for PSMA radioligand therapy underwent [18F]FDG-PET and [68Ga] PSMA-PET with low-dose CT to assess if they were candidates for PSMA radioligand therapy. The studies were performed within 2 months of each other. Images were retrospectively analysed using MIM encore to compare differences tumour bulk volume (TBV) between the [18F]FDG-PET and [68Ga] PSMA-PET/CT which were then correlated with gleason score, PSA level and survival.

### Results

There were twenty men cancer 44-74 (mean age  $66 \pm 8$ ) with adenocarcinoma of the prostate were included, the average gleason score was 8 ( range 7-10). All patients had pelvic lymph node metastasis and widespread skeletal disease involvement on [68Ga] PSMA-PET. In 19/20 patients, no mismatch findings could be identified. One (1) patient (4,5%) with a gleason score of 10 had FDG+/PSMA- lesions in abdominal lymph nodes , which precluded them from [177Lu]-PSMA therapy. This cost R 500 000 worth of [18F]FDG-PET/CT (20 scans) to save R 240 000 worth of [177Lu]-PSMA RLT - assuming that the patient would have received 4 cycles.

### Conclusions

Our analysis showed that FDG+/PSMA-lesions occurred (5%) of patients prior to RLT, and this was in a patient gleason score of 10. Therefore the cost of FDG PET/CT may be justified in patients with high gleason scores. However, there is evidence that RLT still prolongs survival even in patients with low PSMA avidity as compared to best standard of care. Future studies need to look at the cost-effectiveness of RLT in patients with discordant-FDG positive disease with no other therapeutic options.

## Promoting a validation, protocol of radio-HPLC methods for the determination of radiochemical purity for small size gallium-68 labelled peptide derivatives

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**Introduction:** Radiochemical purity is a test performed during the quality control of radiopharmaceuticals to quantify the content of radioactive impurities introduced or formed during labelling. Possible impurities and labelled products are separated using the High-Performance Liquid Chromatography (HPLC) principle whereafter the radioactive nature of the sample can be detected using radio-detectors. Very often due to this radioactive nature, non-active reference standards, if available, are used to establish and validate HPLC separation method which, in most aspects, can be translated to analyses of radioactive components. However, the radioactive properties of these radiopharmaceuticals will require some specific tests or an adjusted setting concerning the validation of some parameters. In this work, we present step by step how validation parameters are investigated (accuracy, linearity, precision, robustness, stability of solutions) applicable to general radio-HPLC analytical methods, on an example of an already approved radiolabeled compound.

**Methodology:** This method is translatable to the determination of radiochemical purity for most small size gallium-68 labelled peptides or peptidomimetics. Validation parameters described are in accordance with European Pharmacopoeia and ICH prescribed acceptance limits. HPLC method was validated and provides evidence on accuracy, repeatability, intermediate precision, suitable system parameters, specificity, and robustness. The limits of detection and quantification for the method are also provided.

**Results:** Validation parameters met all criteria as following: Accuracy 100%±5, repeatability RSD < 2.00%; intermediate precision RSD < 5.00%; theoretical plates N>1000, capacity factor k>2.00 specificity / resolution Rs >1.50, symmetry As =0.8-1.5. Robustness was established for column temperature (±2 degrees Celsius), flow rate (±10%) and mobile phase TFA (±10%). Solution stability was established (short-term integrity and forced degradation/thermal decomposition).

**Conclusion:** A successful validation of radio-HPLC procedure suitable for conjugated peptides to allow gallium-68 radiolabeling is presented. Radiochemical purities >95% can be trustfully measured. While such procedures are often neglected during research, Good Manufacturing Practices and the necessary validations of analytical equipment are non-negotiable when radiopharmaceuticals are prepared for administration to patients.

## The development of lyophilised kit preparations in a research setting: challenges & lessons learnt

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In recent years, there has been a considerable increase in investigations towards the development of lyophilised (cold) kit formulations as an alternative to the automated synthesis of <sup>68</sup>Ga

radiopharmaceuticals. Kit-derived radiopharmaceutical preparation is not only user-friendly, robust and cost-effective but also offers the convenience of in-house preparation using generator produced

radioactivity. It is these advantages that have contributed to the extensive use and success of <sup>99m</sup>Tc based SPECT-radiopharmaceuticals. As this approach involves efficient radiolabelling techniques and

minimal equipment it is now considered to ease the more technical, manual radiosynthesis of <sup>68</sup>Ga-based PET-radiopharmaceuticals. To date, commercially available kit-based <sup>68</sup>Ga-

radiopharmaceuticals coupled with market authorized <sup>68</sup>Ge/<sup>68</sup>Ga-generator systems have already demonstrated feasibility. Cold kits consist of freeze dried non-radioactive substrates and reagents

(biological moiety, chelator-, buffer-, stabilizer-, and bulking agent). The gold standard follows a

“shake-and-shoot” preparation that facilitates one-step “instant” radiolabelling. Producing

lyophilised kits of acceptable quality requires consideration of all aspects of the design and

manufacturing process. Process validation evaluates data gathered over the product’s lifecycle to

confirm that the process reliably generates a product to a determined standard. Sterile filtration,

aseptic dispensing and freeze-drying are the main steps validated in kit production.

Herein we report our experiences in the process validation of the GMP-manufacture of a lyophilised kit for PET imaging. Several kit batches were produced over a few months (different operators) and

analysed using the validated quality control methods for visual inspection, pH (cold/radiolabelled),

loss on drying, mass of pellet, concentration and radiochemical purity/yield. Filters were consistently

integral and did not impact the concentration or kit pH (solution acidity). Batches subjected to

sterility assessment were found to have no bacterial growth. Following visual inspection, the average

yield of kits was ~80 % with minor, major and critical defect numbers within acceptable limits. The

laboratory scale, benchtop freeze dryer used in this study’s setting did not provide a repeatable and

robust production method. The evaluation revealed the impact of limited freeze dryer functionality

whereby shelf temperature cannot be controlled to facilitate pre-freezing and only minimal

adjustments to temperature and vacuum pressure settings are possible. This finding is of particular

importance for buffer solutions with complex freeze-drying properties.

These factors together with the duration of pre-freezing and freeze-drying had a noteworthy

influence on the visual appearance of the kit pellet and the kit pellet moisture – these results are

highly relevant whenever novel PET-radiopharmaceuticals are under consideration for translation to routine manufacture.



## THE ADDED ADVANTAGE OF 18-FLOURO DEOXY GLUCOSE, POSITRON EMISSION TOMOGRAPHY/COMPUTED TOMOGRAPHY IN A SOLITARY BONE SCAN LESION IN BREAST CANCER

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The paper exhibits the added benefit of 18-Flouro de-oxy glucose, positron emission tomography imaging coupled with computed tomography in a patient with a solitary lesion in the spine on initial staging with SPECT/CT imaging.

Background: Bone scan remains the standard of care for breast cancer patient staging. Positron emission tomography together with computed tomography (PET/CT) is extensively used as an imaging tool for staging, re-staging, and treatment response and evaluating of recurrent disease. Breast cancer is characterized into histological sub groups which exists at molecular level based on estrogen and progesterone gene heterogeneity. One of the sub types is luminal B which encompass 15-20% of breast malignancy. Luminal-B tumors are mostly characterized by highly histological grade, aggressive, high proliferative index and associated with worse prognosis. Hann N et al found that on a lesion basis, Wholebody Flouro de-oxy glucose (FDG) PET/CT was more sensitive and equally specific for detection of bone metastases when compared with bone scintigraphy.

Material and methods: A case of a 69 year old female patient with histologically proven luminal B, stage III A left breast cancer, ki-67 of more than 14%. The patient had Tc-99m MDP bone scan and was imaged using whole body scintigraphy, which was subsequently followed by FDG PET/CT.

Results: MDP whole body bone scan imaging showed an isolated intense uptake in the spinous lesion at the level of T1 which was localized in the posterior elements however, due to high intense uptake coupled with relative hypo density on CT, right axillary lymph nodes and high clinical staging; further investigation with 18F FDG PET/CT was done. History of trauma was excluded. Findings on PET/CT showed widespread metastases involving the skeletal and further metastases in the lymphatic and visceral organs.

### Conclusion

Results demonstrate that in a patient with highly clinical and Scintigraphic suspicious lesions, a negative bone scan cannot conclusively exclude metastases. Additional imaging with FDG PET/CT is an essential advantage for adequate management of the patient.

## VALUE OF THE HEALTH ASSESSMENT QUESTIONNAIRE DISABILITY INDEX SCORE IN EVALUATION OF RADIOSYNOVIORTHESIS TREATMENT RESPONSE IN PATIENTS WITH REFRACTORY SYNOVITIS

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### Background

Radiosynoviorthesis (RSO) is a medical technique which utilises the administration of intra-articular radiopharmaceuticals to improve the effects of synovitis in inflammatory and haematological joint diseases. Response to RSO is routinely assessed with a Visual Analogue Scale (VAS). A fifty percent (50%) or more reduction in the pre-treatment VAS value six weeks post-therapy is considered successful. The Health Assessment Questionnaire (HAQ) score is a tool validated in patients with rheumatological diseases and haemophilic arthropathy to assess treatment response to conventional medical therapy. There is limited evidence for the use of the HAQ score to assess response after RSO.

### Aim

To determine the accuracy of the HAQ score in the response assessment of patients treated with radiosynoviorthesis for refractory synovitis.

### Method

A retrospective study of patients with refractory synovitis due to inflammatory or haematological joint disease was conducted. They received radiosynoviorthesis treatment at Groote Schuur Hospital between 2017 and 2023. Active synovitis was confirmed through radiological, sonographic, and/or scintigraphic imaging. Patients were clinically examined and were required to complete HAQ and VAS questionnaires prior to and six weeks after treatment. Patients without both sets of HAQ and VAS scores were excluded from the study.

### Results

Of the thirty-three patients treated and forty-five total treatments performed, twenty-one patients were included in the study, with a combined total of twenty-four treatments. Most patients were excluded due to incomplete HAQ and/or VAS scores. More than half of the patients suffered from rheumatoid arthritis (54%). Haemophilia was the second most common disease (25%). The remainder comprised of patients with psoriatic arthritis, reactive arthritis, and juvenile idiopathic reactive arthritis (20%). There was a sixty three percent (63%) decrease in VAS scores post RSO of fifty percent (50%) or more compared to the pre-treatment scores. A twenty-five percent (25%) improvement in the HAQ scores post RSO treatment was identified. No significant correlation was found between the percentage change in VAS and HAQ scores in patients with successful RSO response.

### Conclusion

There was a significant improvement in the HAQ score of patients receiving radiosynoviorthesis. However, the HAQ score alone cannot be used as an independent assessment tool in the evaluation of treatment response. Integration of the HAQ score with other assessment parameters, including relevant clinical and imaging criteria, is therefore required for accurate assessment of response to RSO.

# RADIOPHARMACEUTICAL INVENTORY MANAGEMENT AT THE NUCLEAR MEDICINE DEPARTMENT OF DR GEORGE MUKHARI ACADEMIC HOSPITAL

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**BACKGROUND AND OBJECTIVES:** Radiopharmaceutical products and procedures are highly expensive, hence there is need to reduce unnecessary costs incurred. An appropriate inventory management system is essential in ensuring traceability of each radiopharmaceutical product, prevention of stock outs and expired products. In this study the inventory management system in place before the appointment of a radiopharmacist at the SPECT Nuclear Medicine Department (DNM) of DGMAH was reviewed. An appropriate inventory management system was recommended, and SOPs and stock card systems were developed.

**METHODS:** The study was descriptive and quantitative. In phase 1, a review of the inventory system utilized in the Department was conducted, baseline findings were examined according to GPP guidelines, and problem areas were detected. In phase 2, the most appropriate reorder quantity, average daily consumption and lead times for each radiopharmaceutical product were calculated.

**RESULTS:** The overall compliance per focus area at baseline (Phase 1) was as follows: inventory management (0%), availability of SOPs (16.7%), record keeping (33.3%), temperature monitoring (33.3%), physical storage (66.7%) and radiopharmacy security (66.7%). In phase 2 the identified non-compliance indicators were rectified and recommendations were made.

**CONCLUSION:** The study revealed that the inventory management system at the DNM was informal. It was characterised by stock outs, expired medication, lack of a stock card system, lack of inventory management SOPs and ineffective record keeping procedures. Interventions were introduced to address these gaps. Thus, an independent demand inventory system was employed by developing stock cards for reordering and appropriate record keeping allowing forecasting of stock requirements. A radiopharmacist was appointed as part of the research recommendations.

# THE POTENTIAL DANGER OF A PERFUSION ONLY SPECT CT STUDY IN THE DIAGNOSIS OF PULMONARY EMBOLISM SINCE THE ADVENT OF COVID-19 INFECTION

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## Introduction

In recent years nuclear medicine imaging of pulmonary embolism has evolved, with VQ SPECT/CT being the latest technique. When for any reason the ventilation part of the study cannot be performed, the perfusion study is read alone with the CT. It is well known from published studies in the literature that a perfusion only study is associated with a significantly lower diagnostic accuracy. In the initial period of the COVID-19 pandemic, a lot of nuclear medicine centers omitted the ventilation part of the study. COVID-19 has been known to be associated with a lot of perfusion abnormalities, some with corresponding mosaic hypoattenuation on the CT, including pulmonary embolism. Some of these abnormalities reported with a CT only can mimic pulmonary embolism and further reduce the diagnostic accuracy of a perfusion SPECT/CT study. In this study we looked at the data of known COVID-19 patients who had perfusion SPECT/CT studies and were erroneously diagnosed as having pulmonary embolism initially. We also looked at a small cohort of patients who performed perfusion only SPECT CT during a period where we could not perform ventilation studies and compared their study to a repeat study performed with a ventilation.

## Methods

This was a retrospective study where we reviewed the perfusion only SPECT/CT studies of 9 recovered COVID-19 patients and 4 patients with unknown post COVID-19 status. We compared these findings with a repeat study which included ventilation studies.

## Results

Thirteen cases were studied in total. All thirteen cases had perfusion only SPECT/CT studies reported as being positive for pulmonary embolism. A review of their repeat studies with ventilation revealed that these defects were matched, and these matched defects were associated with mosaic hypoattenuation on CT. Some of these patients also had these matched defects persisting for over a year.

## Conclusion

The COVID-19 pandemic might contribute to a further reduction in the diagnostic accuracy of a perfusion only SPECT/CT study. Our hypothesis is that perfusion only studies with a defect that corresponds to an area of subtle mosaic hypoattenuation should be at best reported as a non-diagnostic study, with appropriate follow up management. We would however need bigger and well organized prospective studies to validate this.

## 99mTc ETHYLENEDICYSSTEINE-DEOXYGLUCOSE (GLUCOSAMINE) IN THE IDENTIFICATION OF ACTIVE DISEASE IN PATIENTS WITH RHEUMATOID ARTHRITIS – A SINGLE CENTER PROSPECTIVE STUDY

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### Introduction

Rheumatoid arthritis is a chronic inflammatory disease, which, if not treated properly, leads to irreversible joint damage, deformities, and premature mortality. There are several investigations that can be used to detect and assess disease activity in patients with this condition. These include, clinical assessment, laboratory investigations and imaging. However, these different techniques have various shortcomings, and ideally a modality with a very high diagnostic accuracy is needed to detect disease activity, even at a subclinical level. The aim of this investigation was to evaluate the utility of 99mTc-labelled glucosamine (ethylenedicysteine-deoxyglucose) for the identification of active disease in patients with rheumatoid arthritis. The hypothesis is that this radiotracer could be a cost-effective, highly sensitive tool in the assessment of patients with rheumatoid arthritis.

### Methods

This was a prospective study in which 22 participants with diagnosed active rheumatoid arthritis using the American College of Rheumatology/European League Against Rheumatism (ACR/EULAR) were recruited. Glucosamine ligand was synthesized according to standardized existing protocols in the literature described by Yang et al. Each participant was injected with 20-25mCi of 99mTc-labelled glucosamine. Flow, blood pool and SPECT/CT imaging of the most symptomatic joints was performed, in addition to a delayed whole-body imaging 2 hours after radiotracer injection. Joints were qualitatively assessed for abnormal increased uptake of the radiotracer. Degree of uptake was interpreted as normal, mild, moderate or severe using a 4 point scoring system.

### Results

Twenty-two participants were recruited into the study. The median (IQR) age was 59 (49-68) years and majority (95.5%) were females. All 22 participants had abnormal increased uptake of tracer in sites of known disease, including unknown sites. SPECT/CT imaging localized tracer uptake specifically to the synovial space. Increased flow studies were associated with either grade 3 or 2 disease on the delayed static images.

### Conclusion

99mTc-labelled glucosamine was able to clearly detect active disease (synovitis) in all the known symptomatic joints of our patient population. The intensity of tracer uptake seems to be related to disease severity. It is yet to be known if this can also provide prognostic information.

## SPECT/CT whole-body bone scan time optimization using image quality: A phantom study

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'Introduction:

Aim: Investigate the effect of different reconstruction protocols on image quality, when using varying scan times on single-photon emission computed tomography/Computed tomography (SPECT/CT). The main goal was to establish the shortest possible acquisition time for clinical whole-body SPECT/CT whilst maintaining an acceptable image quality.

Methods:

The National Electrical Manufacturers Association (NEMA) image quality phantom from PTW with 5 fillable spheres ( diameters:10, 13, 17, 22 and 28 mm [37mm sphere was unusable]) without the lung insert was used. The Siemens dual head Intevo 6 gamma camera with a Low energy-High resolution (LEHR) collimator was used for the Technitium-99m energy. The spheres were filled with activity making a sphere-to-background concentration ratio of 8:1. Acquisition times were varied with a time per projection (TPP) of 5, 8 and the department's 15 seconds (60 projections per head). These TPP's gave a total of 1 bed acquisition of 5, 8 and 15 minutes. Images were reconstructed on a MIMS software workstation using the ordered subset expectation maximization algorithm (OSEM), with 8 subsets and 4, 8 and 16 iterations with a 3D Gaussian filter smoothing of 0-12 mm at full width half maximum (FWHM). The image quality components that were evaluated were Contrast recovery (CR), signal-to-noise ratio (SNR) and background variability (BV).

Results

It was observed that the image quality statistics improved with increasing acquisition time. In respect of iterations, it was observed that the convergence of CR was observed at 8 iterations, while the higher 16 iterations tended to increase the reconstruction times significantly. The 3D Gaussian filter used for smoothing was found to be decreasing the noise at 12mm FWHM. The BV improved with increased filter smoothing for all sphere sizes with noise decreasing as acquisition time increased. At a time of 8 seconds, the possibility of detecting the smallest sphere from the images was greater than 5 as per the rose score criterion.

Conclusion

This study was aimed at finding the optimal imaging time which can lower/reduce the departments scan times. From this study, we observed that the 8s (8 min per bed) image qualities were similar with our departments 15s (15min per bed) imaging. Our protocol of 3 bed whole body SPECT/CT total time of 45min could be lowered by up to 21min to a total WB scan time of 24min. The optimal reconstruction parameter chosen from the results was 8 subsets, 8 iterations and 12 mm filter.

## VALIDATION OF PLANAR GATED BLOODPOOL LEFT VENTRICULAR EJECTION FRACTION PROCESSING USING MONTE CARLO SIMULATED DIGITAL PATIENT PHANTOMS

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Background: Left ventricular ejection fraction (LVEF) is an important parameter for evaluating cardiac function. Commercial software programs are used to calculate LVEF values from planar gated blood pool (GBP-P) studies acquired with gamma cameras in nuclear medicine. However, before implementing these programs in clinical practice, it is essential to validate the accuracy and precision of the LVEF values obtained with these software programs using phantom studies. The aim of this study was to assess the accuracy of LVEF values obtained with a commercial software program using gamma camera-simulated GBP-P studies of digital patient models.

Methods: The SIMIND Monte Carlo program, along with the 4D-XCAT (XCAT) phantom, was utilised to simulate 69 male and female GBP-P studies with different combinations of LVEF values (19.9% - 77.4%) and end-diastolic volumes (86ml - 155ml). These studies were then converted to DICOM format and transferred to a processing station with the available commercial software program, ALPHA NUCLEAR (AN). The 'known' LVEF of the XCAT phantom, determined from the exact volume of the left ventricle in the end-diastolic and end-systolic positions, was considered the gold standard. Processing was performed by three independent, experienced operators, and LVEF values were reported. Intra- and inter-variability were evaluated using the interclass correlation coefficient (ICC). A regression analysis was performed to compare the real LVEF and processed LVEF values.

Results The intra-observer variability for all three observers was excellent (ICC1 = 0.99; ICC2 = 0.99; ICC3 = 0.98). The ICC evaluating the interobserver variability was 0.98. When comparing the average LVEF obtained by the three observers using the AN software, with the known XCAT LVEF, a good correlation was obtained ( $r^2 = 0.94$ ). A slight underestimation of the XCAT values at lower LVEF values was seen ( $LVEF_{AN} = 1.09 LVEF_{XCAT} - 7.59$ ).

Conclusion Independent processing of GBP-P studies using AN software indicated excellent intra- and interobserver variability. A good agreement was found between the calculated LVEF values using the AN software and the known LVEF of the XCAT simulated images. The digital patient models created with the 4D-XCAT phantom provided a database of studies with known LVEF values whereby AN commercial software was validated. This study will be expanded by evaluating different commercial systems using the same simulated GBP-P studies.

## ESTIMATION OF RADIATION DOSE TO COMMUTERS USING PATIENT TRANSPORT AND THE NURSING STAFF FROM THYROTOXICOSIS PATIENTS TREATED WITH RADIOACTIVE IODINE-131 AT UNIVERSITAS ACADEMIC HOSPITAL.

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**Introduction:** Thyrotoxicosis patients are treated with radioactive I-131 to control the disease. The challenge comes after the I-131 treatment. The patient needs to be accommodated in a single room and separate rooms as per SAHPRA regulations, public dose below 1 mSv/year. The patients treated at Universitas Academic Hospital (UAH) come from the Free State, Eastern Cape, Northern Cape, and Lesotho. Due to prevailing economic disparity after discharge, some patients may thus be obliged to travel long distances (4-7 hrs) in crowded public transportation (e.g. FS Health Patient Transport Services). The ongoing concern of the dose to nursing staff caring for these patients was also addressed.

**Aim:** The study aimed to determine the 24 hrs radiation exposure from thyrotoxicosis patients' post-treatment with I-131.

**Methods and Materials:** Two electronic alarm dosimeter was used to measure radiation exposure in the room of admitted I-131 patient, and A total of 85 patients were monitored (Oct 2021 – Dec 2022). The dosimeter was at the wall (head side of the patient bed), 1.5 m above the floor, and 0.2 m from the patient's head. The second set of measurements was done at the wall (feet side of the patient bed), 1.5 m above the floor, and 2 m from the patient's abdomen. The average time to monitor the room's radiation levels was approximately 18 hours—the mean prescribed I-131 activity was  $15.7 \pm 2.9$  (ranging from 12 mCi to 20 mCi).

**Results and discussion:** The results show that radiation dose in the room varied from 106 to 6220.0  $\mu\text{Sv}$ , (Median: 385.0  $\mu\text{Sv}$ , Mean:  $545.0 \pm 793.0$   $\mu\text{Sv}$ ) for approximately  $18.6 \pm 6.9$  hours post I-131 administration. The results indicate that the I-131 patients might give significantly higher radiation doses to other passengers using patient transport due to the reduced distance between them. The results for the second part show that radiation dose in the room varied from 32.0  $\mu\text{Sv}$  to 149.0  $\mu\text{Sv}$  (Median: 57.0  $\mu\text{Sv}$ , Mean;  $66.8 \pm 35.5$   $\mu\text{Sv}$ ) for approximately  $18.6 \pm 2$  hours after I-131 administration.

**Conclusions:** The SAHPRA regulation of isolating patients after the I-131 therapy capsule ensures that the family and public are safe from radiation exposure. The dose received by the nursing staff is low. However, the dose to passengers in patient transport can be higher if patients are treated as outpatients without considering socioeconomic factors.



## MANAGERS' PERSPECTIVES OF SERVICE DELIVERY IN PUBLIC SECTOR NUCLEAR MEDICINE DEPARTMENTS IN SOUTH AFRICA

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### Introduction

Nuclear Medicine (NM) is a rapidly growing discipline. Across SA, there are 14 and 46 NM Departments in the public sector and private sector respectively. Most SA public sector hospitals are strained due to a lack of adequate resources resulting in delayed bookings and patient waiting lists because of fewer examinations being performed daily. Service contracts between private and public hospitals as a solution for patient access to NM come at a high cost and proves ineffective in curbing the number of unassisted patients.

This research aimed to assess the service delivery of NM Departments in the public healthcare system from the managers' perspectives.

### Methodology

This is qualitative research that employed the exploratory research design. The data was collected using a structured interview mostly conducted telephonically. In this research, 11 out of 14 managers in the 14 NM Departments from five Provinces in SA participated. Interviews were mainly open-ended and centered on addressing shortcomings perceived in ensuring adequate service delivery to public sector NM Departments.

### Results

Three themes pertaining to service delivery in NM emerged, namely: barriers affecting service delivery; enablers that influence service delivery; recommendations to improve service delivery. The majorly devastating findings were on barriers affecting service delivery. These comprised five categories namely: poor service delivery resulting from the unavailability of different resources, poor working conditions, the effects of the COVID-19 pandemic, inadequate management and administration, and limited access to NM Departments, especially in rural communities. Accolades accomplished by different NM Departments were illustrated as enablers. Hard-working departments provided quality NM services for the population and had personnel willing to go the extra mile despite the barriers noted. The recommended solutions were to improve what is available, decentralize services to rural communities, allow personnel who work in the field to implement some of the advances in NM and use of technology and training to improve the services of the NM Departments.

### Conclusion

It was evident from the study that barriers have a devastating impact on service delivery. The unavailability or the lack of key resources was regarded as a major cause of discomfort for healthcare workers and patients. These limitations impact negatively on NM in the clinical setting as a therapeutic department and lead to suboptimal service delivery. There is a lot to be done at all levels of the healthcare system in order to optimize service delivery on NM in the public healthcare sector.

## VALIDATION OF AN IN-HOUSE DEVELOPED THERAPEUTIC DOSIMETRIC SOFTWARE TOOL FOR THE TREATMENT OF <sup>177</sup>LUTETIUM-DOTATATE PEPTIDE RECEPTOR RADIONUCLIDE THERAPY

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Different computational software tools used in nuclear medicine have been written to improve radiation dose assessment, especially in therapeutic nuclear medicine. While many studies have investigated therapeutic nuclear medicine dosimetry, it has been noticed that very few papers compare the therapeutic software tools to each other. The aim of our study was therefore to validate our in-house developed software tool Masterdose using the commercial software OLINDA/EXM 1.0.

Methodology was based on clinical patient data treated for neuroendocrine tumours with <sup>177</sup>Lutetium (Lu)-DOTATATE at a South African hospital. All patients underwent the same SPECT acquisition protocol and were corrected for scatter, partial volume, collimator-detector response, gamma camera calibration and attenuation. Correction factors (CFs) were applied to images to convert counts to activity. The first cycle of peptide receptor radionuclide therapy (PRRT) for 11 single photon emission computed tomography (SPECT) patients were compared on the Masterdose and OLINDA/EXM 1.0 software tools at 1, 24, 72 and 168 hours. Cumulated radionuclide activity and the absorbed dose were compared for the two software tools. The absorbed dose difference was then compared using statistical Bland-Altman analysis.

Masterdose and OLINDA/EXM 1.0 had different peptide receptor radionuclide therapy methodologies. This led to different results obtained for the software tools. The OLINDA/EXM 1.0 software did not propose steps for image quantification. Therefore, Masterdose quantification results were compared to the Dosimetry Toolkit<sup>®</sup> (DTK) on the Xeleris<sup>®</sup> software by GE Healthcare. The largest cumulated activities were found in tumours compared to the kidneys for both software. On average the relative percentage difference (RPD) between the cumulated activities of Masterdose and DTK was 10.5 % and 10.9 % for the kidneys and tumours respectively. Absorbed dose differences were similar in magnitude. For the kidney Bland-Altman analysis, a particular trend was seen, the calculated absorbed doses were almost consistently higher than the bias or systematic error of 0.2 Gy. However, the tumour Bland-Altman analysis show a non-systematic difference, as calculated absorbed doses were distributed evenly around the bias of 1.9 Gy.

Due to the different therapeutic approaches of the two software compared, results obtained during the comparison affected the results of this study. The RPD between the software were attributed to different methodologies during the CFs and activity quantification of Masterdose and OLINDA/EXM 1.0.

## CORONARY [68GA] GA NODAGAZOL PET VERSUS CORONARY ARTERY CALCIUM SCORING IN THE EVALUATION OF ATHEROSCLEROTIC ACTIVITY.

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### Introduction:

Coronary artery calcium (CAC) scoring is a reliable method for cardiovascular risk stratification and predicting outcome in patients with coronary atherosclerotic disease (CAD). [68Ga]Ga-NODAGAZOL PET/CT is a novel molecular technique for imaging microcalcification, a hallmark of atherosclerotic plaque instability. In our previous work, we demonstrated a positive correlation between [68Ga] Ga-NODAGAZOL uptake and the cardiovascular risk profile of patients. The aim of this study was to establish the relationship between [68Ga] Ga-NODAGAZOL uptake and CAC score in atherosclerotic plaque and to assess the cardiovascular outcome.

### Methods:

18 patients with CAD were prospectively recruited to undergo cardiac CT and [68Ga] Ga-NODAGAZOL PET/CT within two weeks of each other. The CAC score for each of the three coronary arteries was computed. The patients were followed-up for major cardiovascular events including death, non-fatal myocardial infarction, or new-onset angina. [68Ga] Ga-NODAGAZOL uptake in the atherosclerotic plaques in the respective arteries was quantified using the maximum standardized uptake (SUVmax). For each artery, the atherosclerotic plaque with the highest SUVmax was used as the representative lesion for the respective artery. The SUVmax was correlated to the CAC score using the non-parametric Spearman-rank test, whereas the Mann-Whitney U test was used to determine the correlation of CAC and [68Ga] Ga-NODAGAZOL uptake with cardiovascular outcome.

### Results:

94% of patients had BMI >25kg/m<sup>2</sup> (94%) whereas 67%; 56%; 72% and 61% had a history of dyslipidaemia, systemic hypertension, diabetes mellitus, and smoking respectively. The patients tolerated the [68Ga] Ga-NODAGAZOL with no untoward side effects. The mean injected activity [68Ga]Ga-NODAGAZOL was 144.67MBq (range =92.5 - 244.2). The median (Range) CAC score was 118.3(1.1-1326.1). The SUVmax of [68Ga] Ga-NODAGAZOL uptake in LAD, RCA, and LCx arterial territory were as follows Median (Range): 2.05(1.56-3.34), 2.05(1.33-3.41)and 1.23(1.06-2.75), respectively. CAC score demonstrated a strong positive correlation with the SUVmax of [68Ga]Ga-NODAGAZOL uptake  $r=0.545$  ( $p < 0.001$ ). CAC score of the different arterial territories also showed significant correlation to cardiovascular outcome (LAD,  $p=0.011$ ; RCA,  $p=0.021$ ; and LCx,  $p=0.013$ ). Similarly, [68Ga] Ga-NODAGAZOL uptake measured using SUVmax showed significant correlation to cardiovascular outcome in the LAD ( $p=0.026$ ) and RCA ( $p=0.021$ ) but not in the LCx ( $p=0.052$ ) arterial territories.

### Conclusion:

There is a strong positive connection between CAC score and the degree of coronary [68Ga] Ga-NODAGAZOL uptake in atherosclerotic plaques in a CAD cohort. Furthermore, both CAC and [68Ga] Ga-NODAGAZOL uptake exhibit a strong positive correlation with long-term cardiovascular outcomes.



## APPLICATION OF MICROPET-CT TO IN-VITRO COMPUTATIONS

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Clinical Positron Emission Tomography-Computed Tomography (PET-CT) is routinely used for cancer staging and therapeutic response. Preclinical evaluation of novel radiopharmaceutical compounds requires in vitro cancer cell studies with PET-CT investigations for imaging of cancer cells that have been attempted<sup>1</sup>. However, higher system resolution is needed for in vitro tests. A microPET-CT ( $\mu$ PET-CT) offers adequate resolution; however, literature lacks on the use of this modality for in vitro application. Acquisition parameters and methodology need to be established to provide a validated method for using  $\mu$ PET-CT when imaging cell plates.

A  $\mu$ PET/CT scanner with associated accessories (carbon fibre plate, <sup>22</sup>Na-point source and post-processing software) along with an automated gamma counter was utilised for the in vitro analysis. Cancer cells were cultivated in a six-well plates to a suitable starting density. <sup>18</sup>F-FDG (1.8MBq $\pm$ 0.3MBq) was administered into each well and was left for a 60 min incubation period. Each well was subjected to three wash cycles using phosphate buffer solution(PBS) to remove the excess media and un-bound <sup>18</sup>F-FDG. Each rinsed solution was counted on the gamma counter and a standard dilution of <sup>18</sup>F-activity was created as a reference for activity quantification. Imaging was commenced immediately following the last wash cycle using  $\mu$ PET/CT. Following the  $\mu$ PET/CT acquisition, the cells were scraped from each well, collected in PBS and the activity was measured using the gamma counter. A field of view containing the 6-well plate was reconstructed and the volume of interests associated with each well was quantified. <sup>18</sup>F-FDG-image guided uptake/ well was verified by sample gamma counting.

The <sup>18</sup>F-FDG- $\mu$ PET/CT imaging of cells was successful, i.e., sufficient dose yielded a good count rate and signal-to-background ratio. Tracer uptake in cancer cells was detected and further used to quantify normalized uptake values (kBq/mL) which correlated well with the <sup>18</sup>F-FDG signal in the cell pellet by using gamma counting. The acquisition method was also optimised over a series of acquisitions to ensure future acquisitions show minimal variances.

The acquisition parameters and methodology for in vivo imaging using  $\mu$ PET/CT were established. Therefore, this study is precedent that the use of  $\mu$ PET/CT imaging is not limited to in vivo procedures but can also be successfully applied to an in vitro setting. This will further streamline the translation of pre-clinical research to the clinical field.

## APPLICATION OF SPECT/CT AND CERENKOV LUMINESCENCE FOR IMAGING OF THERAPEUTIC ISOTOPES

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Therapeutic radionuclides are not easily visualised and are paired with positron ( $\beta^+$ ) or gamma-emitting radionuclides to evaluate disease progression using conventional nuclear imaging techniques (PET; SPECT). Popular theranostic pairs include 68Ga/177Lu, 68Ga/255Ac, 64Cu/67Cu or 188Re/99mTc. SPECT imaging of therapeutic isotopes is not routinely used because of the generally low yield in gamma emissions of these isotopes. However, technological advances in SPECT imaging hardware-and-software have made imaging of these tracers more feasible.

Cerenkov-derived luminescence imaging (CLI) detects light emitted because of charged particles from radioactive-decay moving faster than light in a medium; therefore, CLI is suitable to directly visualise beta-emitting therapeutic radionuclides. This aspect should be utilized more to support clinically relevant questions. This study evaluated SPECT-and CLI-capabilities of three, relatively unused, therapeutic nuclides (186Re, 188Re, 109Pd) in comparison to already evaluated therapeutic nuclides (177Lu,131I) and diagnostic (68Ga, 64Cu, 18F, 89Zr) nuclides for application of theranostic radiopharmaceuticals.

SPECT/CT - each isotope was calibrated according to manufacturer specifications using a multi-pinhole collimator. Images for all radionuclides were acquired using Derenzo and PET NEMA phantoms. Following image reconstruction, the images were evaluated according to: sensitivity, resolution, uniformity and contrast signal-to-noise-ratio (SNR).

CLI - A 1:2 dilution series of each radionuclide (1 mL in saline) was pipetted into a 24-well plate (activity 0–9.16 MBq/mL). Each radionuclide underwent CLI at half-life appropriate time points. Radionuclides were compared with respect to activity-concentration (kBq/ $\mu$ L) and average radiance (p/s/cm<sup>2</sup>/sr).

SPECT/CT - Isotopes with gamma rays in the vicinity of 140keV showed the best sensitivity and resolution (188Re and 186Re); visual uniformity of Rhenium-188 was superior to Rhenium-186. Compared to 177Lu and 131I evaluation of the SNR of 186Re- and 188Re-isotopes and 109Pd correlated as expected.

CLI - Linear regression and range analysis for all nuclides indicated a strong, positive correlation between the light emission intensity and activity concentration ( $R = 0.99$ ). High energy emissions of 188Re ( $E_{\beta\max} = 2.10$  MeV;  $E_{\beta(\text{ave})} = 0.8$  MeV; 70% intensity) corresponded to radiance observed for 68Ga ( $E_{\beta\max} = 1.89$  MeV;  $E_{\beta(\text{ave})} = 0.8$  MeV; 87% intensity) while radiances for 109Pd and 186Re were 2.5- and 5-fold lower, respectively (all other nuclides  $\leq 15$ -fold lower).

Rhenium-186/-188 exhibits superior SPECT imaging capabilities over 131I- and 177Lu-radionuclides while 109Pd still provides suitable SPECT images. All three therapeutic isotopes have excellent CLI properties comparable to other therapeutic-/PET-isotopes. The SPECT- and-CLI imaging capabilities of these therapeutic radionuclides allow for further research into more comprehensive targeting mechanism.

## PRECLINICAL PET/CT AS IMAGING MODALITY TO SHOWCASE THE EFFECT OF CHEMICAL PROPERTIES ON OVERALL IN VIVO BIODISTRIBUTION.

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Radiopharmaceuticals for diagnostic imaging through positron emission tomography (PET/CT imaging) are designed to target specific diseases. The design of these radiopharmaceuticals during the drug development process focusses on chemical features such as stereochemistry, steric hindrance, molecular modelling and structure-activity relationship. However, when it comes to application of these compounds in vivo, there are additional factors that affect the compound's biological distribution, excretion and targeting efficacies which need to be considered<sup>1</sup>. These chemical properties include: molecular weight, lipophilicity and overall charge (if applicable) of a compound.

The aim of this retrospective study was to compare the microPET/CT images of 7 selected radiopharmaceuticals and correlate the visualised biodistribution to the drugs chemical properties.

The radiopharmaceuticals were formulated in saline or 0.1M PBS with pH 7-7.4 and administered intravenously into the mouse tail vein (volume < 200 µL and activity dose of 8-10 MBq per animal). Mice were imaged over 20-30 min in list mode using microPET/CT. Timepoints for imaging were 1h, 2h, 4h, and 24h dependant on the radionuclide used and its corresponding half-life. Images were reconstructed iteratively with attenuation and activity correction to allow for quantification. The visualised biodistribution and excretion of the following radiopharmaceuticals: <sup>18</sup>F-FDG, <sup>68</sup>Ga-PSMA-11, <sup>64</sup>Cu-TETA, <sup>64</sup>Cu-GluCAB2-NH<sub>2</sub>, <sup>89</sup>Zr-DFO-SC, <sup>64</sup>Cu-GluCAB1-Mal-HAS and <sup>89</sup>Zr-DFO-IIIB6 were compared, and the results correlated to the compounds molecular weight, lipophilicity and overall charge.

The metabolism and excretion of the radiopharmaceuticals follows either a renal (kidney) or hepatobiliary (liver, intestines) pathway. Smaller compounds (under glomerular filtration size) that are generally negatively charged with a less lipophilic structure follow a renal excretion route while larger compounds (over glomerular filtration size) that are often protein/antibody bound, generally positively charged and more lipophilic, exhibit slow clearance through a hepatobiliary pathway. Exceptions were noted when desired biodistribution effect based on compound size were overshadowed by the charge and lipophilicity of the compound.

Different radiopharmaceuticals present with different biodistribution and excretion patterns that can be effectively visualised using microPET/CT. The chemical properties (molecular weight, nett charge and lipophilicity) of a radiopharmaceutical affect the biodistribution and subsequent excretion pathway and therefore need to be considered when designing new radiopharmaceuticals.

## SCROTAL WALL PERTECHNETATE WASHOUT AS A PREDICTOR OF TESTICULAR SALVAGE FOR MISSED TORSION BY TESTICULAR SCINTIGRAPHY

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**BACKGROUND AND OBJECTIVE:** Testicular torsion (TT) is a urological emergency, early diagnosis and intervention prevents long term complications including male infertility. In our case series we aimed to distinguish if the washout of tracer from the scrotum could be predictive of salvageability of the testes in patients with testicular scintigraphy (TS) diagnostic of testicular torsion.

**METHODS:** We evaluated the TS scans of patients that underwent surgical intervention for orchidectomy for suspected missed TT from January 2016 to December 2021 at Dr George Mukhari Academic Hospital. We assessed if the washout of tracer in the scrotum between 5-minute and 20-minute images and determined the correlation with salvageability of the testes in patients scintigraphically diagnosed with missed TT. Semiquantitative (maximum and minimum counts) and visual analysis was conducted.

**RESULTS:** A total of 12 patients underwent surgery. Of these, 50% of the testes were salvaged and in 50%, the testes could not be salvaged and complete orchidectomy was performed. On visual and semiquantitative analysis, we detected a progressive decrease in pertechnetate uptake in the scrotal wall uptake in the salvaged testes. The maximum percentage of decrease in counts in this group was 34%, the median time to presentation after the onset of symptoms was 2.5 days with an interquartile range of 2 days and a range of 1 day to 5 days. In the non-salvaged group, visual and semiquantitative increase in uptake in scrotal wall was noted. The maximum percentage increase in counts was 32%. The median time to presentation after the onset of symptoms in these patients was 6 days with an interquartile range of 5 days and a range of 2 days to 14 days

**CONCLUSION:** This case series demonstrates an association between pertechnetate washout and salvageability of the testes. Testicular salvage is noted in patients who demonstrated an progressive decrease in uptake of pertechnetate in the scrotal wall over a 20 minute imaging time in patients who presented beyond the golden 6 hour time limit. TS may be a great tool in identifying those patients with higher possibility of salvage and assist to prioritize their surgery.



## SYSTEMATIC ASSESSMENT OF ANAESTHETIC PROCEDURES FOR BETTER ALIGNMENT AND COMPLIANCE WITH RADIOTRACER MICROPET-CT IMAGING IN SMALL ANIMALS

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Non-invasive, functional imaging techniques such as Positron Emission Tomography-Computed Tomography (PET/CT) provide the means for studying physiological and pharmacological processes in real-time, both in animals or humans 1. PET/CT imaging has made a major footprint in preclinical research of pharmacology, molecular biology and medical sciences, facilitating innovation in neuroscience, cancer, inflammation and cardiovascular research. Rodent animal models are commonly anaesthetised with ketamine or isoflurane in order to prevent movement artefacts during image acquisitions 2; however, such anaesthesia often alters the physiological condition of the rodents. Consequently, the microPET/CT findings obtained in anaesthetized animals may not accurately represent the physiological properties 3. This investigation aims to systematically align different anaesthetics with common radiotracers for PET/CT imaging studies to provide information that can enhance the radiotracer representation according to their targeting mechanism in rodents to advance the quality and translatability of microPET/CT-derived data.

The following compatible anaesthetics were mainly assessed for suitability of using [18F]FDG-, but also for [68Ga]Ga-DOTA-tate-, or [68Ga]Ga-PSMA-11-microPET-CT diagnostics: isoflurane, sevoflurane, pentobarbital, propofol, or combination of fentanylcitrate-fluanisone/diazepam, and ketamine/xylazine. These anaesthetic agents were compared based on animal physiology (respiration rate, body temperature, apparent glucose metabolism and expected radiotracer biodistribution).

The investigated anaesthetics caused altered respiration rate, body temperature, glucose metabolism and have effects on target organs which affected radiotracer biodistribution. Isoflurane inhalation is the most desired anesthetic procedure; however, both sevoflurane and isoflurane (also ketamine/xylazine administration) induced hypoglycaemia. Sevoflurane can be suggested over isoflurane for cardiac imaging. Propofol is suitable for [18F]FDG imaging but should be avoided for neuroimaging causing decreased cerebral glucose metabolism. Pentobarbital does not alter the blood glucose metabolism but has a lower safety margin for rodent anaesthesia. Fentanylcitrate-fluanisone/diazepam the preferred injectable anaesthetic option for [18F]FDG being the least interfering with glucose metabolism. Tumor imaging quality using [68Ga]Ga-DOTA-tate- or [68Ga]Ga-PSMA-11-microPET/CT was supported by isoflurane or sevoflurane; however, higher glucocorticoid levels and certain neuroendocrine abnormalities occurred (may originated by hypothermia).

This investigation proved that anaesthetics used for preclinical research using microPET/CT imaging need to be carefully aligned with the radiotracer's half-life, and metabolism, to warrant an uninterrupted image acquisition of high quality. A better understanding of the interplay between radiotracers and available anaesthesia agents can allow for the best choice of anaesthesia that will improve the performance and outcome of imaging studies. Thus, radiotracers are supported to show their full research potential to become non-invasive in vivo biomarkers or sensitive drug efficacy monitoring tools.

## MANAGEMENT OF PEDIATRIC DTC INCLUDING SERBIAN EXPERIENCE

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Juvenile differentiated thyroid carcinoma (DTC) is a rare neoplasm, accounting for 0.5%–3% of all childhood malignancies. According to the Serbian Cancer Registry 42 newly diagnosed DTC ≥19 years were reported during the period 1999–2010; female-to-male ratio of 3.7:1. Juvenile DTC has aggressive clinical appearance showing cervical nodal involvement in most of the cases, often associated with distant metastatic disease. However, comparing with adults, the overall mortality rate in children is not increased. Guideline for juvenile DTC treatment has been published yet. Thyroid cancer in children and adolescents is usually treated empirically. I-131 activity can be adjusted by body weight (1.85–7.4 MBq/kg); by age (1/3 of the adult activity is administered to a 5-year-old; 1/2 of the adult activity in a 10-year-old; and 5/6 of the adult activity in a 15-year-old); by surface area, or by 24-h thyroid bed uptake body weight (<5% uptake would receive 50 MBq/kg; 5–10% uptake would receive 25 MBq/kg; and 10–20% uptake would receive 15 MBq/kg). In some institutions, empiric (fixed) activities of 1.1–11.0 GBq are applied. Dosimetry is another option which enables I-131 treatment on a calculated patient-specific basis. In patients with extensive lung metastases, one must consider possible high-absorbed dose in lungs and consequently recommend lower radioiodine activity. In 2015, The American Thyroid Association Guidelines Task Force on Pediatric Thyroid Cancer published the Management Guidelines for Children with Thyroid Nodules and Differentiated Thyroid Cancer with new recommendations. In our institution, we analyzed 51 juvenile DTC with median follow-up of 10 years. Eleven patients developed recurrent disease. The probability of recurrence was 16.7% at 5 years, 22.3% at 10 years, and 33.3% at 15 and 23 years after the initial treatment. Strong predictive factors for recurrence were age ( $p < 0.001$ ), initial treatment ( $p < 0.0001$ ), and tumor multifocality ( $p < 0.011$ ), while sex, nodal metastases at presentation, distal metastases at presentation, histologic type of the tumor, T stage, and clinical stage had no influence on relapse. The outcome of disease at the last check-up was excellent: 90.2% patients achieved CR, 3.92% patients achieved PR, while 1.96% patients had stable disease. Two patients have died (one disease-related death). Conclusions: dosimetry should be performed in patients with extensive lung metastases, and if retained radioiodine activity in the lung exceeds 3 GBq, the I-131 activity should be reduced to avoid the risk of radioiodine-induced lung fibrosis. Treatment with radioiodine is safe, with no adverse effects on fertility or secondary malignancy.

## INDIVIDUALIZED CALIBRATION OF ESTIMATED GLOMERULAR FILTRATION RATE FOR SERIAL RENAL FUNCTION MONITORING

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**Background:** Glomerular filtration rate (GFR) is widely accepted as the best index of renal function. Estimated GFR (eGFR) calculated by serum creatinine remains the most widely used method in clinical practice. However, eGFR lacks precision and accuracy in early stages of kidney disease (stages 1 and 2) and is not a reliable measure of change in GFR over time. Therefore, in certain groups of patients where decisions based on eGFR may have significant clinical consequence, measurement of GFR is recommended. However, the currently used methods to measure GFR are comparatively expensive and time consuming and are not widely available. As a compromise between measured and estimated GFR, particularly for patients who require serial GFR measurements, it is possible to develop individualized eGFR equations by calibrating the first GFR estimate using an initial GFR measurement taken at the same time. By applying an individualized calibration factor to subsequent estimates, it may be possible to provide an improved individualized estimate of renal function. This would be beneficial in settings where there is limited access to GFR measurement.

**Methods:** Patients who had undergone two radionuclide based GFR measurements within 3 months of serum creatinine measurement at the nuclear medicine division, Tygerberg Hospital, between January 2009 and December 2019 will be retrospectively identified. The 2021 CKD-EPI Creatinine equation will be used to calculate the estimated GFR. A patient specific calibration factor (Kpt) will be calculated from the initial measured (mGFR1) and initial estimated GFR (eGFR1), where  $mGFR1 = Kpt * eGFR1$ , i.e.  $Kpt = mGFR1 / eGFR1$ . For subsequent studies a patient specific eGFR (eGFR-Pti) will be calculated using the patient specific calibration factor and the estimated GFR at that particular time ( $eGFRi = Kpt * eGFRi$ ). Bland-Altman analyses will be used to compare eGFR and patient specific eGFR to measured GFR.

**Results:** Selected patients will be described in terms of their demographic profile, clinical indication, and relative timing of creatinine and GFR measurements. The results of Bland-Altman analyses the performance of patient specific eGFR (eGFR-Pti) to conventional eGFR in terms of bias and limits of agreement.

**Conclusion:** Individualized calibrated GFR is expected to perform better than estimated GFR in serial renal function monitoring. This would provide a practical alternative to serial GFR measurement in environments where there is limited access to GFR measurement.

## SHOULD FLUORODEOXYGLUCOSE POSITRON EMISSION TOMOGRAPHY INTEGRATED WITH COMPUTED TOMOGRAPHY REPLACE METHYLENE DIPHOSPHONATE BONE SCINTIGRAPHY IN THE MANAGEMENT OF BREAST CANCER PATIENTS WITH SUSPECTED SKELETAL METASTASES?

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Breast cancer is the leading cause of cancer related death in women and the fifth leading cause of mortality worldwide. Bone metastases is the most frequent site of dissemination. Prevention of morbidity from metastases to bone requires accurate, early diagnosis for preliminary staging, planning of treatment, treatment monitoring, restaging and prediction of survival in patients with breast cancer. Many centers<sup>1</sup> in South Africa still mostly make use of 99m Tc-MDP bone scintigraphy in the staging work-up of these patients, likely due to the relatively low cost, availability, and physician familiarity with this imaging modality. 18F-FDG PET/CT scans are more sensitive in detecting lytic lesions and purely marrow involvement. Additional soft tissue involvement can also be detected using this imaging modality. The purpose of this cross-sectional study is to compare the detection of suspected metastatic skeletal lesions in both 99m Tc -MDP bone scintigraphy and 18F-FDG PET/CT scans as well as its impact on management.

### METHODS:

Forty-six (46) breast cancer patients with a mean age of 52 years, underwent a 18F-FDG PET/CT scan and a 99m Tc-MDP bone scintigraphy performed within 31 days between January 2017 and March 2023. Comparison was done based on a lesion-by-lesion analysis. The results of the image interpretation were compared retrospectively. Any change in stage and management was recorded.

### RESULTS:

A total of 142 lesions were detected on 99m Tc-MDP bone scintigraphy and 237 lesions were detected on 18F-FDG PET/CT scans which led to a change in management of 12 patients with 99m Tc-MDP bone scintigraphy and 17 patients with 18F-FDG PET/CT scans. Of the forty-six (46) patients, 76% (n=35) of patients had invasive ductal carcinoma (IDC), 13% (n=6) had invasive breast cancer no specific type 4(IBC-NST), 4.3% (n=2) had invasive lobular cancer (ILC) and 6.5% (n=3) of patients had missing histological results. In 1/46 (2%), an osteoblastic lesion was detected on the 99m Tc-MDP bone scintigraphy and negative on 18F-FDG PET/CT scan which resulted in change in management. This patient had triple positive IDC. With regards to tumor stage(T), 63% (n=29) had T4 lesions, 17% (n=8) had T3 lesions, 15% (n=7) had T2 lesions and 4% (n=2) had T1 lesions. Detailed statistical analysis to follow.

### CONCLUSION:

18F-FDG PET/CT scans proved to be the superior modality in detecting skeletal metastases and other soft tissue distant metastases resulting in a significant number of patients having a change in management in comparison to 99m Tc-MDP bone scintigraphy.

## PERCEPTIONS OF NUCLEAR MEDICINE TECHNOLOGISTS ON PET/CT IN GAUTENG PROVINCE, SOUTH AFRICA.

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In South Africa, the first positron emission tomography/computed tomography (PET/CT) imaging modality was installed in Gauteng in 2005 with subsequent installations. The introduction of PET/CT requires appropriate staff training, redesign of patient workflow, new skills, problem-solving abilities, and radiation protection. With the introduction of PET/CT in the UK, nuclear medicine technologists (NMTs) encountered challenges in defining their roles, unfamiliarity with the new technology, and new working procedures, which led to confusion and concerns about the new technology.

Since the introduction of the first PET/CT in South Africa, the perceptions of NMTs of this dual imaging technique have not been determined. Therefore, this research study aimed to explore and describe the perceptions of NMTs regarding PET/CT as an imaging modality in nuclear medicine. The objectives formulated to achieve the aim were to establish how NMTs in Gauteng perceive PET/CT as an imaging modality and to make recommendations on managing the needs and desires of NMTs based on study findings.

A qualitative, descriptive phenomenological research design was applied with semi-structured interviews until data saturation was reached. Purposive sampling was employed to select 18 practicing NMTs in Gauteng to participate in this study. The participating NMTs consisted of three cohorts, those with PET/CT experience, those with no PET/CT experience, and those with previous PET/CT experience but not exposed to PET/CT at the time of data collection. Thematic analysis was used to facilitate the management of codes, categories, and themes during the analysis of the data.

The three themes that emerged from the categories are NMTs' perspectives of PET/CT in nuclear medicine, patient management in PET/CT, and PET/CT challenges. The findings suggest that NMTs in Gauteng perceived PET/CT as the future of nuclear medicine. They acknowledged the positive role played by PET/CT in patient management, especially oncology patients. Findings also reveal a gap in PET/CT training in Gauteng, which the NMTs perceived as hindering their career growth in nuclear medicine. NMTs raised concerns regarding high radiation exposure associated with PET/CT imaging, lack of psychological support, single qualification in nuclear medicine, and lack of PET/CT exposure. Based on the study findings, recommendations are made to manage the needs and desires of NMTs in Gauteng. Recommendation for PET/CT training collaboration between the government and private nuclear medicine departments to facilitate CPD-accredited rotation clubs between nuclear medicine departments.

## ASSESSMENT OF THE LONGITUDINAL RADIOLABELING PERFORMANCE OF KIT BASED GALLIUM 68 PROSTATE SPECIFIC MEMBRANE ANTIGEN 11 PREPARATIONS TO SUPPORT THERANOSTICS OF PROSTATE CANCER

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**Introduction:** Prostate-specific membrane antigen (PSMA) is overexpressed in prostate cancer (PC) and has been discovered as a target for theranostics in Nuclear Medicine. Therapy response can be monitored using PET/CT imaging using [68Ga]Ga-PSMA-11, a peptide ligand which can be conveniently prepared from freeze-dried kit starting material currently developed at the South African Nuclear Energy Corporation. This study aimed investigating kit-based [68Ga]Ga-PSMA-11 preparations from a large sample size (n>350). Variations observed within radiolabeling efficiencies, radiochemical purities and yields provided the input assessing the longitudinal radiolabeling robustness and helped determining those parameters and factors leading to lower radiopharmaceutical yields or preparation failures.

**Methods:** An assessment of 388 [68Ga]Ga-PSMA-11 preparations over a one-year period using kit starting material was conducted as a retrospective and cross-sectional study. In total 330 data sets (85.0%) were enrolled in the analysis of the robustness of this labelling technique and also to identify the factors that may cause variations and failure of [68Ga]Ga-PSMA-11 preparations. Systematic evaluation of the responsible issues was addressed to enhance future radiopharmaceutical preparations.

**Results:** A total 196/330 (59.4%) successful [68Ga]Ga-PSMA-11 preparations were achieved (i.e., a radiochemical purity  $\geq 95\%$ ); of those a 52.1% instant preparation (no cartridge purification required) and 7.3% failed rate occurred. For 134 (40.6%) preparations with a radiochemical purities 51-95% and for those yielding 45-50%, the purification process restored appropriate [68Ga]Ga-PSMA-11 product for 73.1% and 11.2% of the preparations, respectively. The level of training to operating personal correlated with the preparation success rate. The ideal radiolabelling reaction pH was 4.5, otherwise causing decrease in labeling efficiency. Adsorbed radioactivity to cartridge material was the main culprit for low radiochemical yields.

**Conclusion:** The performance of [68Ga]Ga-PSMA-11 preparation from kit starting material is easily applicable as a method with a low-cost technique that yields a high-quality product, ideal to be used in radiopharmacy environments with limited resources. It was proven that for most preparations with compromised purity, the product quality was restored by a Sep-Pak cartridge based purification. Future [68Ga]Ga-PSMA-11 preparation success may be further increased by generator quality management, attention to expertise, and correct pH and Sep-Pak cartridge adjustments.

## 177Lu-PSMA Radioligand Therapy in Chemotherapy-naïve Patients, Initial Experience

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Prostate cancer remains a leading cause of mortality despite multiple therapy options. Peptide radioligand therapy (PRLT) with <sup>177</sup>Lutettium-prostate specific membrane antigen (<sup>177</sup>Lu-PSMA) prolongs survival and improve quality of life in heavily pre-treated patients with metastatic castrate resistant prostate cancer (mCRPC). Earlier introduction of PRLT in chemotherapy-naïve mCRPC patients are still not extensively explored.

### Aim

To report initial experience of PSA response and progression free survival (PFS) in chemotherapy-naïve mCRPC patients treated with <sup>177</sup>Lu-PSMA.

### Methods

We retrospectively reviewed 18 chemotherapy-naïve patients with mCRPC who presented for PRLT. The patients received <sup>177</sup>Lu-PSMA at 6-8 weeks intervals with an average activity 8 GBq per cycle. PSA was measured prior each cycle and monitored three monthly after completion of PRLT. PSA response classified as  $\geq 50\%$  decline, any decline and disease progression ( $\geq 25\%$  increase above baseline).

### Results

All patients had metastases on <sup>68</sup>Ga-PSMA PET/CT. Seven and 4 patients had nodal and skeletal metastases only respectively, 6 had both and 1 had additional lung metastases. All patients had good functional status with Eastern Cooperative Oncology Group score of 0 to 1. All patients had received hormonal therapy, 4 had prostatectomy and 7 had pelvic radiation. A total of 90 cycles (median 4, range 2-6) were administered. Following the first cycle, decline of  $\geq 50\%$  in PSA was noted in 22% and any decline in 61% of patients. Seven patients' (39%) PSA increased after one cycle. After the last cycle all 13 (72%) patients who had a PSA response demonstrated decline of  $\geq 50\%$ . Ninety percent of patients who had any PSA decline after the initial cycle had durable PSA response of  $\geq 50\%$ . Fifty-seven percent of patients who had an initial increase in PSA progressed. All patients had an follow-up of at least 4 months. The median PFS was 9 months (range 4-36 months).

### Conclusions

A positive response was noted in this chemotherapy-naïve cohort with durable PSA decline in over 70% of the patients and median PFS of 9 months after <sup>177</sup>Lu-PSMA therapy.

## CHANGING THE MILK SCAN PROTOCOL, PARADOXICAL OR?

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**Background/Aim:** Gastro-oesophageal reflux (GER) and pulmonary aspiration occur in children of all ages. Milk scans have been used for the detection of GER for decades. There are various documented protocols for milk scans in search of GER with no universally accepted protocol. We recently amended our protocol from imaging patients for 30 minutes and assessing gastric emptying at 2 hours to imaging patients for 60 minutes and assessing gastric emptying at 3 hours due to the potential of missing episodes of reflux and overclassifying delayed emptying when normal. This study aims to assess the value of the change in protocol in being able to detect significant number of refluxes.

### Method:

We retrospectively reviewed the scans of all patients who presented for milk scans at the Red Cross War Memorial Children's Hospital from 1 November 2021 to 30 November 2022. The number of reflux episodes detected in patients imaged with the 30-minute protocol in comparison to the number detected with the 60-minute protocol was reviewed. The episodes of reflux were classified with an in-house severity scale and gastric emptying was evaluated at different time intervals. Delayed gastric emptying at 2 hours was quantified as residual activity in the stomach >37% and at 3 hours as >80% emptying into the bowel.

### Results:

A total of 200 studies were reviewed over the 13 month period (103 males and 97 females).

In the 30-minute studies, 400 reflux episodes were observed in 70 patients, 7 were classified as normal, 9 mild reflux, 19 moderate reflux, 35 severe reflux, with 21 patients demonstrating delayed gastric emptying at the 2-hour images. 11/21 patients with delayed gastric emptying had no reflux and 10/21 patients had reflux of varying classifications.

In the 60-minute studies, 537 reflux episodes were observed in 65 patients, 16 were classified as normal, 8 mild reflux, 15 moderate reflux, 26 severe reflux with 12 patients demonstrating delayed gastric emptying at the 3-hour images. 8/12 patients with delayed emptying had no reflux and 4/12 had reflux of varying classifications.

### Conclusion:

More episodes of reflux were detected in the 60-minute reflux search without increasing incidence in patient numbers. Our findings did not demonstrate substantial differences owing to a change in the protocol. A standardized study comparing similar patient populations with the question of reflux may be beneficial in order to establish a common practice.



## EMERGING RADIOISOTOPES; THE PRODUCTION THEREOF AND DECAY PRODUCT CONSIDERATIONS

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There are a number of exciting new radioisotopes that are proposed, evaluated and produced in research quantities all claiming to be the best new radioisotopes for the future. Different aspects are highlighted; Decay mode (alpha, beta, Auger), half-life, radiochemistry, production routes to name a few. In this paper radioisotopes will be evaluated in the light of the above but also in the particular South African setting with the view to the future MPR (Multi-purpose Reactor) Necsa is planning to build to replace SAFARI-1 in the early 2030's.

Several of the new radioisotopes proposed do not consist of single decay and some have several daughters contributing to the radiation/therapeutic dose – also referred to as in vivo generators. The underlining physics and radiochemistry aspects of these will be presented and what it means for the chelators employed in the targeting vectors.

## DE NOVO NUCLEAR MEDICINE SERVICES IN THE PUBLIC HEALTH SECTOR OF NORTH WEST PROVINCE: TURNING VISION INTO REALITY

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Klerksdorp/Tshepong (K/T) Hospital complex is a provincial tertiary hospital that serves the entire North West Province in South Africa. Commencement and expansion of hospital-based tertiary services in this health facility has been on-going for the past two decades with many successes. Non-availability of Nuclear Medicine services in the public health sector of this province have resulted in patients being referred to academic hospitals in Gauteng with its attendant logistic and cost-related burden.

The journey of the first Nuclear Medicine clinical unit in the North West province public sector from conceptualization to realization culminated in its completion within the projected financial year. The implementation process had inevitable minor adaptations expected for a project of such complexity. However, this is an accomplishment that was made possible through strategic planning, creative problem-solving, and resourceful partnerships with relevant stakeholders. Funding was secured from NHI Oncology grant, equipment sourced and installed, trained staff recruited, and clinical service commenced, ensuring that the long-term vision turned into reality.

The establishment of Nuclear Medicine at K/T hospital complex has been a significant achievement that will cater to the needs of the North West patients, enhancing diagnostic and therapeutic services by increasing capacity and volume of services rendered, and ultimately improving overall patient outcomes.

Our success has improved referral processes and provided quality services at an affordable cost, reducing the burden on tertiary hospitals in Gauteng. Partnership with Wits Nuclear Medicine in Johannesburg provides flexibility and ease of referral pathway as we continue to expand our range of services.

In conclusion, starting Nuclear Medicine services in the North West province public health sector has been an overall success story. Our experience demonstrates the importance of foresight, strategic planning and partnerships in turning vision into reality.

By establishing the first nuclear medicine unit in the public health sector of the North West Province, we have improved accessibility and provided a higher quality of care to patients in the region. We hope to inspire similar efforts to expand nuclear medicine services in other relatively underserved regions.

## The SA experience of Re-SCT production

### Sello T

Rhenium-188 is a beta emitter with a half-life of 17 hours and is able to penetrate the human tissue to a depth of 2-3 mm. This makes this therapeutic isotope ideal for the treatment of superficial tissue with no risk to other parts of the body. The relatively short half-life allows treatments to be performed in a very short period of time. Re-SCT is a specialized non-invasive brachytherapy for the treatment of basal and squamous cell carcinomas of the skin. Application of this epidermal radioisotope therapy facilitates local direct cell-killing as a result of the beta-radiation. In this process both the local death of cells and the local reactions of the immune system of the body to repair itself is triggered.

In partnership with the company currently supplying SCT to several parts of the world, doses were manufactured at Necca and distributed to various hospitals as part of a market penetration study. The recipient hospitals would have access to the specially designed equipment that is needed to measure and apply the final dose for application. The production of the Re-SCT involved the elution of a Tungsten-188 (W-188) generator with saline into the production unit. Re-188 was processed via a chemical reaction to afford the end product. The paint based product was transferred into Re-SCT carpoules which were packaged and sent out for patient treatment. As the first such study in the country, we report the technical challenges and triumphs experienced in the installation of the production set up in our existing infrastructure. Even though a lot of challenges were met, several doses were successfully prepared and used in patient treatment.