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Radiological imaging in cystic fibrosis: cumulative effective dose & changing trends over 2 decades

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8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25	 O'Connell OJ, MD, Cork Cystic Fibrosis Center, Cork University Hospital, University College Cork, Cork. McWilliams S, MD, Cork Cystic Fibrosis Center, Cork University Hospital, University College Cork, Cork. McGarrigle AM, PhD, Dept. of Radiation Physics, Cork University Hospital, University College Cork, Cork. O'Connor OJ, MD, Dept. of Radiology, Cork University Hospital, University College Cork, Cork. Shanahan F, MD, BSC, FRCPI, FRCP (UK), FACP, FRCP(C), Dept. of Medicine, Cork University Hospital, University College Cork, Cork. Mullane D, MD, Cork Cystic Fibrosis Center, Cork University Hospital, University College Cork, Cork. Eustace J, MB, MRCPI, MHS, MD, Dept. of Renal Medicine, Cork University Hospital, University College Cork, Cork. Maher MM, MD, FRCSI, FFR RCSI, FRCR, Dept. of Radiology, Cork University Hospital, University College Cork, Cork. Plant BJ, MRCPI, MFSEM, MD, Cork Cystic Fibrosis Center, Cork University Hospital, University College Cork, Cork.
26 27 28 29 30 31 32 33 34 35 36 37	Guarantor: Dr Barry Plant, Consultant Pulmonologist, Director of Cork Adult Cystic Fibrosis Center, Cork University Hospital, University College Cork, Cork. Ireland. Barry.Plant@hse.ie Study Design: OJOC, SMcW, OJOC, DM, MMM, BJP Statistical Analysis: SMcW, JE Manuscipt preparation: OJOC, AMcG, FS, MMM, BJP

Abstract

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Objective:

- 3 With the increasing life expectancy for patients with cystic fibrosis (CF), and a known
- 4 predisposition to certain cancers, cumulative radiation exposure from radiological imaging is of
- 5 increasing significance. This study explores the estimated cumulative effective radiation dose
- 6 over a 17 year period from radiological procedures, and changing trends of imaging modalities
- 7 over this period.
- 8 **Methods:** Estimated cumulative effective dose (CED) from all thoracic and extra-thoracic
- 9 imaging modalities and interventional radiology procedures for both adult and pediatric CF
- patients, exclusively attending a nationally designated CF center between 1992-2009 for
- 11 >1year, was determined. The study period was divided into 3 equal tertiles and estimated CED
- 12 attributable to all radiological procedures was estimated for each tertile.
- 13 **Results:** 230 patients met inclusion criteria (2,240 person-years of follow-up; 5596 radiological
- procedures). CED was >75mSv for 1 patient (0.43%), 36 patients (15.6%) had a CED between 20-
- 15 75mSv, 56 patients (24.3%) had a CED between 5-20mSv and in 138 patients (60%) the CED was
- estimated to be between 0-5mSv over the study period. The mean annual CED/patient
- increased consecutively from 0.39mSv/yr to 0.47mSv/yr to 1.67mSv/yr, over the tertiles 1-3 of
- the study period respectively (p<0.001). Thoracic imaging accounted for 46.9% of the total CED
- and abdomino-pelvic imaging accounted for 42.9% of the CED respectively. There was an
- associated 5.9 fold increase in the use of all CT scanning per patient (p<0.001).

- 1 **Conclusion:** This study highlights the increasing exposure to ionizing radiation to CF patients as
- 2 a result of diagnostic imaging, primarily attributable to CT scanning. Increased awareness of
- 3 CED and strategies to reduce this exposure are needed.
- 4 MeSH Terms (3-5): Cystic Fibrosis [C08.381.187], Radiation Dosage [N06.850.810.250],
- 5 Diagnostic Imaging [E01.370.350]

6 **Introduction**:

- 7 Experimental and epidemiological evidence has linked the cumulative exposure to ionizing
- 8 radiation, even at low doses, with an age-dependent increased risk for the development of
- 9 malignancy ¹⁻⁴. A number of recent studies have highlighted the issue of cumulative radiation
- 10 exposure towards hospitalized patients^{2,3}. However, few studies have addressed this issue in
- 11 patients with chronic diseases and there are no studies to date quantifying the annual
- 12 cumulative exposure to ionizing radiation incurred as a result of diagnostic and interventional
- radiology procedures for patients with cystic fibrosis (CF). With the progressive improvements
- in life expectancy for patients with CF, the cumulative exposure to ionizing radiation
- attributable to diagnostic imaging and interventional radiology procedures is of increasing
- 16 significance⁵⁻⁷.
- 17 A number of recent studies have highlighted that CF patients have an increased standardized
- incidence ratio (SIR; the ratio of the observed to the expected number of cancers in the cohort)
- 19 of developing a large number of malignancies, with notable increases in the SIR for thoracic
- cancers, cancers of the digestive tract (SIR 5.6), kidney (SIR 14.0), thyroid (SIR 9.8) and
- 21 lymphoma (SIR 7.3)⁸. The issue of exposure to ionizing radiation in the diagnostic range in this

cohort raises additional concerns. Linking the exposure of iatrogenic ionizing radiation to 1 2 malignancy is controversial, however data within the general population has approximately linked each 10 millisieverts (mSv) of radiation exposure to an incremental mean increase risk in 3 cancer of 1 radiation induced malignancy per 1000 patients, although this risk is significantly 4 affected by both age of exposure and gender 9-10. Secondly, previous studies have proposed 5 that additional caution is required in patient sub-groups (e.g. Crohn's disease patients) who 6 7 because of their chronic relapsing illnesses require repeated diagnostic imaging throughout 8 their lifetime and who already have an increased lifetime risk of developing cancer, may suffer as a result of possible synergy between these two factors resulting in an ever higher incidence 9 of developing cancer ¹¹. 10 11 The benefits of radiological imaging are well proven and have been comprehensively characterized for CF patients 12-14. In CF patients, thoracic CT can detect lung changes from early 12 infancy and before they become apparent by pulmonary function testing ¹⁵⁻¹⁶. An increasing 13 number of clinical trials are using thoracic CT scores as surrogate end points in their studies ¹⁷⁻ 14 ¹⁸. Additionally, some centers use high resolution CT in the routine monitoring of disease 15 progression for pediatric patients with CF¹⁹. 16 This is the first study to examine the cumulative effective radiation dose (CED) incurred by a 17 cohort of adult and pediatric patients with CF, and the changing trends in use of radiological 18 19 imaging modalities in this cohort, over 17 years. It is also the first study to characterize relative 20 anatomical radiation distribution including the percentage of cumulative exposure arising from

- 1 thoracic and abdomino-pelvic imaging and the relationship between clinical phenotype and
- 2 CED.

3 Methods:

4 Study Population:

- 5 All pediatric and adult patients with CF who exclusively attended a nationally designated Adult
- or Pediatric CF center, for all medical therapy for > 1 year from the 1st July 1992 to the time of
- 7 lung transplantation, hospital treatment transfer or death until the study end date of 1st May,
- 8 2009 were included in this study. Ethical approval for the study was granted by the institution's
- 9 research ethics committee. Approval number (reference number) for this study: ECM
- 4(z)03/03/09. All demographic, clinical data and hospital attendances were obtained by chart
- 11 review. Lung function was measured in accordance with the standards set out by the European
- 12 Respiratory Society and recorded as maximum annual FEV₁% predicted using the European
- 13 Community for Coal and Steel (ECCS) reference values²⁰.

14 Data Source:

- 15 Details of all imaging studies and interventional radiology procedures performed on adult
- 16 (>18yrs) and pediatric (<18yrs) patients in the study cohort over the 17 year study period 1st
- July 1992 to 1st May 2009 were obtained from the computerized radiology information system
- of the radiology department. All studies were requested for clinical purposes only, by
- 19 experienced hospital clinicians, predominantly by attending pediatricians and attending
- 20 pulmonologists with a special interest in cystic fibrosis. The CED was measured in millisieverts
- 21 (mSv), a measure designed to represent the overall detrimental effect of a non-uniform ionizing

radiation exposure and useful for population-level comparisons across different types of 1 radiation exposure ²¹. As per previous studies ^{11 22}, estimations were based on a recent article 2 by Mettler at al, which provided a compilation of the mean effective doses for radiological and 3 nuclear medicine examinations from the recent published literature, across the USA, Canada, 4 Japan, Australia and Western Europe, over a similar time period as this study²³. In 5 circumstances where this source was insufficient, estimates were calculated from other 6 7 published sources or extrapolated from doses reported for similar procedures relevant to our studied population over a similar time period and using similar dose calculation methodology ²⁴. 8 9 CED was determined for general radiography (Gen), computed tomography (CT), general interventional procedures (GIV), nuclear medicine (NM), fluoroscopy (FI) including all barium 10 examinations (Ba), and data were then subcategorized into the anatomical area imaged as 11 thoracic, abdomino-pelvic and other. All computed tomography (CT) scans during the study 12 13 period (1992-2009) were performed using either an incremental protocol on either a single slice CT (Siemens Medical Solutions, Erlangen Germany) or a four slice detector CT (Toshiba Aquilon, 14 Toshiba Medical Systems, Zoetermeer, The Netherlands). 15 16 Pediatric effective doses for chest radiographs, abdominal films and barium studies were calculated using the presets for pediatric patients aged 1, 5, 10 and 15 years in the software 17 package PCXMC (version 1.5)²⁵ in conjunction with local hospital exposure parameters and 18 19 protocols. Typified normalized effective doses for pediatric CT scanning were calculated based on validated anatomically-specific Monte Carlo phantom simulations for 1, 5, 10 and 15 year 20 olds²³ (table 1). 21

Statistical Analysis:

- 2 The study period was divided into three tertiles of equal duration from 1st July 1992- 28th
- 3 February 1998 (Tertile 1), 1st March 1998-30th November 2003 (Tertile 2), and 1st December
- 4 2003-1st May 2009 (Tertile 3). Data compilation and statistical analyses were performed using
- 5 Microsoft Access 2007 (Microsoft Corporation, Washington, USA) and SPSS version 15 (SPSS Inc,
- 6 Chicago IL, USA). Algorithms calculating the age of each patient at time of scanning and
- 7 adjusting to the closest categorical age of 1, 5, 10, 15 years old or adult as appropriate were
- 8 established and the person years of follow up in each tertile determined. Comparison between
- 9 groups was performed using Mann Whitney U tests or Pearson's Chi-squared for categorical
- 10 variables. Spearman analysis was used for comparisons of non-parametric data where
- 11 appropriate. Comparison across 3 or more groups was performed using ANOVA or Kruskal
- 12 Wallis tests in accordance with their distribution. A type I error rate ≤ 0.05 was considered
- 13 significant.

Results:

- 15 Study population: 230 patients met inclusion criteria and were included in the final analysis.
- The mean age of patients at the end of tertile 3 was 21.5 years (SD +/-11.6); mean FEV₁ %
- 17 predicted was 65.8% (SD +/- 27.3%). 7 patients received lung transplantations during the study
- period and 42 patients died during the study period, 1 from hepatocellular carcinoma.
- 19 Imaging Modality:

- 1 There were 5596 radiological procedures: General radiographs n= 4730, Ultrasonography n=
- 2 406, CT n= 241, interventional procedures n=127, fluoroscopy n=74 and nuclear medicine n=18,
- 3 over the 2,240 person years of follow up.
- 4 Cumulative Effective Dose:
- 5 Over the total study period 1 patient (0.43%) had a CED >75mSv, 6 patients (2.6%) had a CED
- 6 50-75mSV, 30 patients (13%) had a CED 20-50mSv, 26 patients (11.3%) had a CED 10-20mSv, 29
- 7 patients (12.6%) had a CED 5-10mSV and 138 patients (60%) had a CED between 0-5mSv. In
- 8 total 63/230 (27.4%) of patients had a CED >10mSv over the study period. Plain radiographs
- 9 account for 74% of the total number of studies, resulting in 6% of total cumulative radiation
- 10 exposure for the most recent tertile (tertile 3, see figure 1). Conversely, CT accounted for 8% of
- total number of studies whilst resulting in 74.8% of total radiation exposure in tertile 3 (Figure
- 1). A breakdown analysis of the 10% of patients with the highest CED showed 61.9% related to
- 13 CT, 10.4% from fluoroscopy, 12.7% from general radiography and 15.0% from interventional
- 14 procedures.

- 16 Changing trends over time:
- 17 The mean number of radiological procedures for tertile 1-3 was 2.89 procedures per person per
- 18 year for tertile 1, 2.7 procedures per person per year for tertile 2 and 2.71 procedures per
- 19 person per year for tertile 3. The equivalent annual mean effective dose (aMED) was 0.39 mSv
- 20 per person per year for tertile 1, 0.47mSv per person per year for tertile 2 and 1.67mSv per
- 21 person per year for tertile 3 (p<0.001) (table 2). The increased aMED predominantly related to

- a 5.9 fold increase in the number of CT studies over this time (p<0.001). Similar trends were
- 2 seen in the pediatrics' subgroup analysis (table2).
- 3 Correlation between clinical phenotype and CED: CED correlated with age (r=0.307; p<0.001)
- 4 and in subgroup analysis of adult patients at the end of tertile 3, CED correlated inversely with
- 5 FEV₁% predicted (r=0.182; p<0.04). There was no correlation between mean annual effective
- 6 dose and gender (p=0.125) or cystic fibrosis class mutation (p=0.122).

Anatomical Area Imaged:

- 8 Cumulative radiation exposure, for the whole population, from thoracic imaging, as a
- 9 percentage of total CED was 46.9% over the 3 tertiles with relative exposure of 36%; 34.4% and
- 10 51.7% for tertiles 1-3 respectively, correlating with the increased use of CT thorax. Of the total
- 11 radiation exposure to the thorax, 76.4% (748mSv) was from CT thorax (n=130 scans of thoracic
- region), 9.5% (71.3mSv) from plain radiography, 0.8% (7.93mSv) from fluoroscopy, and 15.5%
- 13 (152mSv) from interventional procedures (figure 2)
- 14 Cumulative radiation exposure, for the whole population, from abdominal imaging as a
- 15 percentage of total CED was 42.7% over the 3 tertiles with cumulative exposure per tertile of
- 16 50%; 45.3% and 41% for tertiles 1-3 respectively reflecting both an increased use of abdomino-
- 17 pelvic CT imaging with a decreasing use in fluoroscopy. Overall the CED to the abdomino-pelvic
- area increased significantly across the 3 tertiles, however there was a relative decrease in the
- 19 percentage contribution from abdomino-pelvic imaging due to the more significant increase in
- 20 thoracic imaging over the same time period. Of the total radiation exposure to the abdomino-

- 1 pelvic region; 53.5% was from abdomino-pelvic CT imaging, 11.9% from plain radiography,
- 2 25.6% from fluoroscopy, and 9.0% from interventional procedures.

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Discussion:

- This is the first study to calculate annual CED from all diagnostic imaging and interventional radiology procedures in patients with cystic fibrosis. 27.3% of our patients had an estimated CED exposure of >10mSv over the study period. Although controversial, an exposure of 10mSv of radiation dose has been predicted to result on average in 1 radiation induced malignancy per 1000 patients¹⁰. Recent studies have highlighted the increased risk for certain malignancies in patients with CF⁸. Several epidemiological studies have shown direct evidence of increased cancer-related mortality following long-term exposure to low levels of ionizing radiation, including diagnostic radiation ^{10 26}, and that this cancer risk follows a "linear no threshold" risk, indicating all radiation exposure may pose a risk of developing cancer²⁷. Studies have suggested that the majority of physicians are not aware of the effective radiation dose associated with radiological procedures, nor do they discuss the risks and benefits of CT and other imaging examinations with their patients ²⁸. One study found 75% of radiologists and physicians significantly underestimated the radiation dose from a CT scan ²⁸. Our paper address's these issues and generates a summary of effective doses for common adult and pediatric imaging modalities, which clinicians can apply to their practice.
- The present study demonstrates a significant increase in the annual mean effective dose to our patients over the past 17 years, with a mean annual exposure of 1.67mSv per patient in the

most recent tertile. This dose may appear relatively modest, given for example a background 1 environmental radiation exposure annually of ~3.9mSv/yr amongst the Irish population²⁹, 2 however this represents a significant exposure for such a young cohort of patients, made all the 3 more pertinent given the progressive increased life expectancy for these patients with 4 improved management of cystic fibrosis⁴. Over the 3 tertiles the frequency of medical imaging 5 6 remained relatively constant; however there was a significant shift in the use of imaging 7 modalities. The single largest contributing factor to the increased radiation exposure was the 5.9 fold increase in all CT imaging (thoracic, abdomino-pelvic and other) from the 1st tertile to 8 the 3rd tertile. This is consistent with international data on the rapidly increasing availability 9 and utilization of CT imaging in clinical medicine ³⁰. 10 11 The changing trends in radiation exposure held similar results for both adult and pediatric populations. The pediatric effective doses we established, are frequently much larger than 12 adult doses (table 1), this is due to the thinner torso in children providing less shielding of 13 organs from the radiation exposure¹. In CT, based on the exposure factors used and in the 14 15 absence of tube current modulation, there is a decrease in effective dose with increasing age. 16 This relates to the increased dose administered per unit of body mass for children as highlighted recently in the "Image Wisely" campaign of the joint North American task force on 17 adult radiation protection³¹. Plain film radiography, however allows for greater dose reduction 18 19 in children, as the exposure parameters required are significantly lower than those used in CT. Increased radiosensitivity in children still exists but the radiation dose decreases with 20 decreasing age, resulting overall in a decreasing effective dose³². The increasing use of CT 21 22 imaging as a routine procedure in some centers and the recent studies which suggest that HRCT

findings frequently pre-date changes in lung function and chest radiograph for pediatric 1 2 patients, highlights the need to develop further strategies to minimize the risks of radiation exposure to these patients³³. Children, particularly females, are inherently more sensitive to 3 4 radiation exposure. This radiosensitivity relates to a large proportion of dividing cells and a longer time for a potential cancer to develop¹. De Gonzalez et al estimated the lifetime risk of 5 radiation induced cancer for CF patients from a modern HRCT thorax protocol at 2 years of age 6 7 to be 24 per 100,000 for females versus 6 per 100,000 males, reducing to 1 per 100,000 for a 50 year old female and 0.3 per 100,000 for a 50 year old male³⁴. The studies to date predicting 8 radiation induced cancer risk from annual thoracic CT for CF patients³⁴ have assumed the 9 cancer rates to be the same as the general population, however recent data supports an 10 increased standardized incidence risk ratios for all cancers in CF patients, suggesting these 11 patients may have an intrinsic increased risk of malignancy. For anatomical reasons, the 12 13 thyroid radiation exposure is higher in pediatric patients undergoing chest radiographs and thoracic CT imaging and the routine use of thyroid and breast shields should be seriously 14 considered in this cohort of patients ³⁵. 15 Despite the fact that greater than 90% of CF patients die from respiratory failure, this study 16 17 highlights that over half of all radiation exposure is related to extra thoracic imaging modalities, 18 contrary to what one might have expected. CF is a multi-system disorder and patients with 19 cystic fibrosis are particular prone to both primary and secondary gastrointestinal disturbances including constipation, distal intestinal obstruction syndrome, acute and chronic pancreatitis 20 and nutritional deficiency. These conditions frequently require diagnostic imaging, which is 21 often of a significantly higher radiation dose than thoracic imaging³⁶. In our cohort of patients, 22

42.7% of radiological imaging was directed at the abdomino-pelvic region. There are consistent 1 2 reports of an increased standardized incidence ratio for digestive tract and other abdominal malignancies in CF patients compared to the general population ^{8 37}. This malignancy risk is 3 likely to be multifactorial, potentially as a result of the inflammation associated with chronic 4 pancreatitis ³⁸, low serum vitamin D levels and an increasing frequency of lung transplantation 5 with associated immunosuppression³⁹. The high abdomino-pelvic radiation exposure is likely to 6 7 pose an additional risk factor for these malignancies. There are a number of available strategies for reducing radiation exposure in patients with CF⁴⁰. 8 9 Patients with CF frequently have a smaller body habitus and reduced body mass indices, so specific low-dose CF protocols optimizing specific parameters such as the millamperes (mA) to 10 patient weight need to be considered⁴¹. Reducing the mAs from 180 to 45 for a conventional 11 thoracic HRCT scan can result in a 4 fold reduction in radiation exposure without any significant 12 difference in image quality ⁴². Additionally, over 97% of CT Thorax scans have significant supra-13 apical and infra-pulmonary unnecessary imaging, resulting in increased exposures to the 14 thyroid and abdominal regions ⁴³. For incremental CT thorax imaging in CF patients, the validity 15 16 of reducing the number of images per CT examination has generated conflicting results, one 17 study has shown an equivalent score using CT cuts from 6 pre-selected sections in a cohort of children with CF 44, whilst others report a reduced sensitivity to detect changes when the image 18 interval is >10mm in children with CF ⁴⁵. With proper attention to detail of scanning 19 parameters radiation dose can be reduced substantially compared to routine chest CT. 20 Strategies currently available for optimization of CT radiation dose include the use of automatic 21 22 tube current modulation, iterative reconstruction techniques and noise reduction filters which

- can lead to significant reduction in radiation exposure without significant impact on image
- 2 quality²⁷. As a result of the preliminary results of this study, we recently implemented a new
- 3 thoracic low dose thin slice CT protocol for pediatric patients with CF which achieved significant
- 4 reductions in mean effective doses¹⁴. Close attention needs to be given to the information
- 5 gained from each imaging request to ensure that the examination is indicated and could not be
- 6 replaced by an imaging study which does not expose the patient to ionizing radiation (e.g.
- 7 ultrasound or MRI). Also it is very important to limit exposure to extra-thoracic organs
- 8 including the development of low dose abdomino-pelvic CT protocols.
- 9 The advantage of detecting early changes on CT imaging awaits additional confirmation, some
- authorities feel strongly that the structural information gained by the use of CT scanning helps
- to tailor treatments, reducing under and over treatment for patients, but to date there are no
- studies proving such a benefit ^{13 19}. The current guidelines suggest there is insufficient evidence
- to recommend use of chest CT scans for routine surveillance but suggest chest CT scanning may
- 14 be helpful in infants with symptoms or signs of lung disease who fail to respond to basic
- interventions⁴⁶. Future studies are also needed to delineate the role of novel strategies in
- thoracic imaging, including the use of hyperpolarized helium magnetic resonance imaging 47
- and positron emission tomography⁴⁰.
- This study has a number of limitations, notably it represents the radiation exposure in a
- 19 single tertiary referral cystic fibrosis center. However, the use of a nationally designated CF
- 20 center acting as both the primary and secondary care facility, regardless of health insurance
- status, ensured the accuracy of capturing of all imaging studies for this cohort of patients. We

acknowledge that retrospective radiation dose exposures appropriate for the period of study 1 involved were used. With technological advances it has been suggested that a reduction in 2 radiation exposure of over 75% can be safely achieved using more modern low-dose CT thorax 3 protocols ⁴⁸, however this may be offset with the increasing use of spiral CT scanning and 4 potentially by the recent suggestions to monitor disease progression using combined PET/CT 5 imaging⁴⁹⁻⁵⁰. The modern use of helical CT scanning in CF patients⁵¹ has increased the diagnostic 6 sensitivity in detecting peripheral thoracic changes³³ but incurs significantly higher radiation 7 8 exposures compared to standard incremental scans of up to 3 mSv per CT thorax. Inevitably there will be variation in clinical practice between centers', one previous French study 9 identified a mean cumulative effective dose from CT scans in CF patients of 19.5 mSv ⁵², with a 10 mean patient exposure to 3.4 CT scans. The annual radiation exposure in our cohort of patients 11 with CF is relatively low, with mean annual exposures of 1.667mSv in the most recent tertile 3, 12 13 this compares to an annual exposure dose of 8.1mSv in a cohort of patient with Crohn's disease at the same institute. It is important to note that only 1 patient (0.43%) had an exposure of 14 >75mSv over the 17 yr period, compared to 15.5% of Crohn's patients¹⁵ and 13% of 15 hemodialysis patients having an exposure of >75mSv over the same time period at our 16 institute¹⁶. However, the critical difference is the young age patients with CF are exposed to 17 18 radiation and the chronic progressive nature of CF from early childhood. Our center follows the 3 fundamental principles of radiation protection; justification, dose optimization through the 19 "as low as reasonably achievable" (ALARA) principle and dose limitation, as set out in the 20 International Commission of Radiological Protection ⁵³.. As a result of this, we have not yet 21 adopted the use of routine CT scanning into clinical practice given the absence of any proven 22

benefit in clinical outcome for such a measure⁵². This study is therefore likely to represent a 1 2 conservative estimate for the annual cumulative effective dose for radiological imaging in CF patients and for the changing trends in CT imaging over the past 17 years. Urgent consideration 3 should be given to the development of low-dose imaging protocols and to regular monitoring 4 5 and recording of CED for patients, particularly in identifiable groups where exposure levels may become high, as in our cohort and in many chronic diseases. Strategies to prospectively 6 7 monitor cumulative dose may include recommendations such as: the creation of dose 8 registries; the mandatory recording of dose within the examination images or report, recording 9 of dose within the patient's medical record and mandatory accreditation of imaging facilities. As a result of this study we are currently implementing a policy of recording the effective dose 10 through the picture archive communication system (PACS) at our institute, to allow for ongoing 11 monitoring and auditing of radiation exposure with each imaging examination⁵⁴. 12 13 **Conclusion:** Patients with CF are exposed to high radiation doses from a young age, exacerbated by the 14 15 increasing use in CT imaging. Strategies need to be developed and implemented with regard to radiation exposure reduction for both thoracic and extra-thoracic imaging in this cohort of 16 17 patients. **Disclosures:** 18 19 There are no competing interests for this paper. The authors grant exclusive publishing rights 20 to the journal, and full ethics board approval was obtained as outlined in the body of the paper. 21 References:

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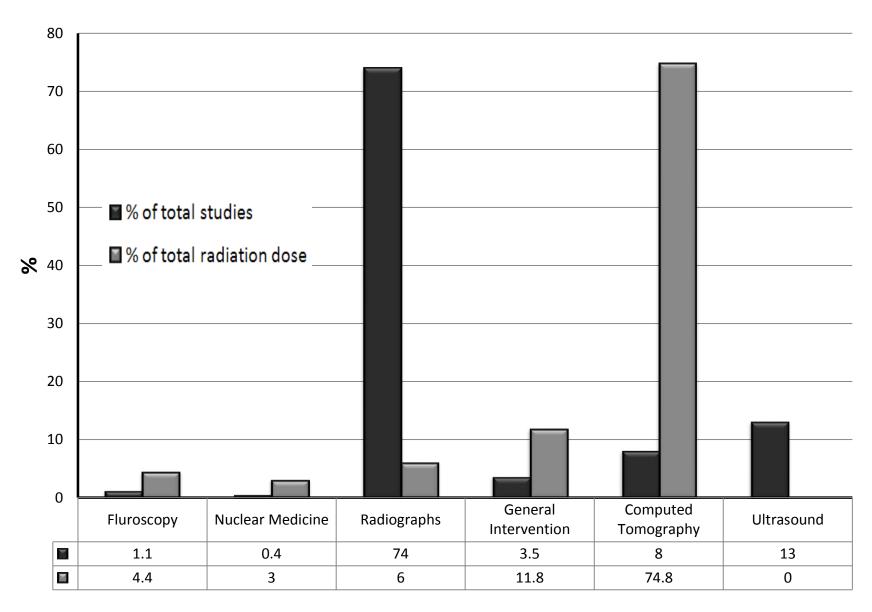
Legend for Table 1: The effective doses of frequent medical imaging procedures used for CFpatients.

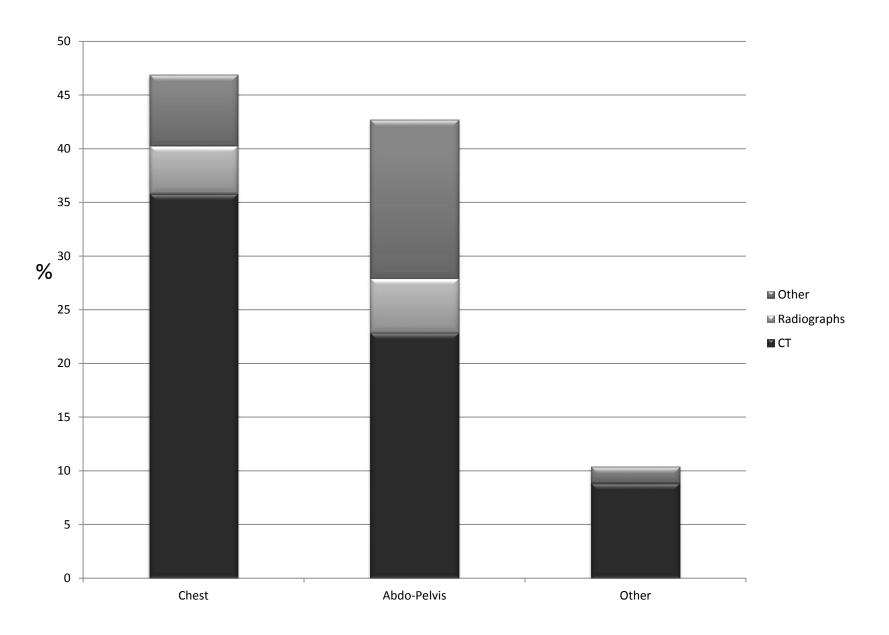
- 40 Legend for Table 2: Total radiation dose per patient year (PPY) across the 3 tertiles for all
- 41 study subjects and secondly for pediatric patients only.

- 1 Legend for Figure 1: Relative contribution of each radiological imaging modality to cumulative
- 2 radiation exposure for tertile 3.
- 3 Legend for Figure 2: Regional percentage breakdown of total radiation dose exposure to
- 4 thorax, abdomino-pelvic and other body parts over tertiles 1-3 for the studied population
- 5 (adult and pediatric inclusive).

Procedure	1yr	5yr	10yr	15yr	≥18yr
	(mSv)	(mSv)	(mSv)	(mSv)	(mSv)
CXR PA	0.003	0.005	0.01	0.014	0.02
Radiograph of abdomen	0.02	0.04	0.1	0.14	0.23
Conventional CT Thorax	13.3	11.2	9.8	7.7	7.0
HRCT Thorax	2.85	2.4	2.1	1.65	1.5
CT abdomen	16	12.8	12	8.8	8
CT Pulmonary Angiogram	28.5	24	21	18	15
Barium Meal	1.28	0.82	0.93	1.63	2.32
Barium Swallow	1.16	0.78	1.09	0.75	0.77
Radiograph Lumbar Spine					1.5

Population	All subjects				Pediatrics only					
Tertile	1st	2nd	3rd	p-value	1st	2nd	3rd	p-value		
Number of subjects	132	157	199		105	115	115			
Cumulative Follow Up										
(years)	599.8	702.6	938.0		435.83	452.17	470.67			
All investigations										
Number PPY	2.89	2.70	2.71	0.86	2.82	2.49	2.04	0.11		
Total dose PPY, (mSv)	0.39	0.47	1.667	<0.001	0.403	0.303	0.788	0.01		
Chest Radiographs only										
No. PPY	2.07	1.93	1.66	0.31	1.98	1.76	1.35	0.03		
Total dose PPY, (mSv)	0.03	0.03	0.03	0.92	0.02	0.02	0.02	0.22		
Abdominal Radiographs Only										
No. PPY	0.23	0.24	0.19	0.81	0.23	0.27	0.07	0.11		
Total dose PPY, (mSv)	0.03	0.03	0.04	0.76	0.03	0.03	0.008	0.07		
			Tota	I CT		_		_		
No. PPY	0.035	0.046	0.210	<0.001	0.042	0.033	0.108	0.006		
Total dose PPY (mSv)	0.136	0.179	1.27	<0.001	0.192	0.084	0.581	0.002		
			Thoracic	CTs only		_		_		
No. PPY	0.015	0.018	0.125	<0.001	0.024	0.018	0.063	0.01		
Total dose PPY, (mSv)	0.087	0.061	0.778	<0.001	0.146	0.045	0.327	0.02		
			ı	elvic CT On		1	ı	1		
No. of PPY	0.001	0.010	0.058	<0.001	0.002	0.002	0.028	0.001		
Total dose PPY, (mSv)	0.008	0.082	0.444	<0.001	0.010	0.012	0.218	0.001		
		T	ı	r CTs	T	1	T	1		
No. PPY	0.019	0.018	0.027	0.65	0.017	0.014	0.018	0.94		
Total dose PPY (mSv)	0.041	0.036	0.054	0.68	0.036	0.027	0.036	0.92		
			ı	tional stud	ı	1	T	1		
No. of PPY	0.034	0.039	0.101	0.01	0.018	0.016	0.033	0.55		
Total dose PPY (mSv)	0.038	0.069	0.172	0.06	0.019	0.008	0.086	0.06		
Total Barium Studies										
No. PPY	0.048	0.039	0.014	0.01	0.048	0.048	0.012	0.03		
Total dose PPY (mSv)	0.104	0.093	0.055	0.36	0.096	0.107	0.026	0.05		
Total Nuclear Medicine Studies										
No. PPY	0.001	0.009	0.011	0.29	0.002	0.008	0.006	0.37		
Total dose PPY (mSv)	0.009	0.051	0.063	0.37	0.011	0.038	0.034	0.49		





Radiological imaging in cystic fibrosis: cumulative effective dose & changing trends over 2 decades

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