

PET-CT in the Pregnant Patient

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Why Consider PET-CT in the Pregnant Patient?

Based on potential risks of fetal loss, teratogenicity, fetal growth retardation and carcinogenesis, positron emission tomography/computed tomography (PET-CT) with an 18F-2-fluoro-2-deoxy-D-glucose injection (FDG) is generally contraindicated in the pregnant patient. Rarely, PET-CT may be selectively performed as a problem-solver in specific pregnant patients. PET-CT is more likely to be performed inadvertently in patients with unsuspected pregnancy. This would most often occur in very early pregnancy, perhaps after only a few weeks gestation, especially at a time that routine urine pregnancy exams are less reliable. Occasionally, human error or miscommunication may be responsible for such events. Medical physicists, radiologists and nuclear medicine physicians may be consulted to evaluate a case and consult with a pregnant patient before or after a planned or unplanned PET-CT examination. 1.2,3,4

Fetal Radiation Associated With the CT Component of PET-CT

For head, neck or chest CT examinations where the fetus is not directly irradiated, maternal Compton scatter radiation doses to the fetus are on the order of 0.2 mGy or less^{5,6}, depending on fetal size and proximity to the primary x-ray beam. With typical abdominal-pelvic CT, the fetus is directly exposed to the scanning beam, with estimated doses of 10- 25 mGy^{5,6} expected. When a risk-benefit analysis in a given clinical situation favors imaging using CT, the goal is to produce diagnostic-quality images with the minimum fetal radiation exposure. For abdominal-pelvic CT in a pregnant patient, depending on patient size, 120 kVp or less and a pitch of greater than 1 may be prescribed.^{5,6} Tube current modulation may be used with care to avoid low noise level settings that could result in a higher fetal dose. Where available, iterative reconstruction methods should be used to improve image reconstruction quality. Multiphase imaging should be avoided.

Normalized fetal CT Monte Carlo dose estimates range from 7.3 to 14.3 mGy/100 mAs⁷. Lazarus et. al.⁸ reported a mean dose of 17.1 mGy (range 8–44 mGy). In a recent series of 54 patients, investigators estimated mean fetal dose to be 24.8 mGy (range, 6.7–56 mGy).⁵

Doses to the fetus from a single-acquisition abdominal-pelvic CT examination have ranged between 10 and 50 mGy in phantom and clinical studies. The most important stochastic risk in this dose range is a potential 1.5- to 2.0-fold increased risk of childhood leukemia. This translates into a relative leukemia risk of 3 in 1000 at background radiation levels to 4 in 1000 at 10 mGy and 6 in 1000 at 50 mGy.⁶



The risks of fetal adverse outcomes, including childhood cancer induction, are small at a dose of 100 mGy and negligible at doses of less than 50 mGy.^{6,7,8}

Use of Iodinated Contrast Agent During Pregnancy

Iodinated contrast agents are known to cross the human placenta and enter the fetus. 11

While iodinated contrast agents have not been associated with teratogenic effects, and there are no reports of clinical sequelae induced by iodinated contrast agents administered IV¹², iodinated contrast agents can cause neonatal hypothyroidism if directly instilled into the amniotic fluid. A recent study of the effect of in utero exposure to a single high dose of iodinated contrast material on neonatal thyroid function identified no significant risks. The <u>ACR Manual on Contrast Media</u> recommends that iodinated contrast agents should be used only as needed in pregnant patients. ¹⁴

Fetal Radiation Associated With the ¹⁸F-FDG Component of PET-CT

Radiopharmaceuticals deliver fetal radiation by two mechanisms (Figure 1 and Table 1):

- Irradiation from radioactivity in maternal organs including bladder, placenta and bowel
- Internal irradiation (beta+ and gamma) from trans-placental radiopharmaceuticals

Typically, smaller moieties of 500 Daltons or less readily traverse the placental barrier. As PET tracers typically represent small molecules, the available PET radiopharmaceuticals ¹⁸F-FDG, Na¹⁸F, ⁸²Rb and ¹⁸F- Florbetapir would be expected to cross the placenta and directly irradiate the fetus. ^{15,16,17,18}

Table 1 - Fetal dose estimated from maternal ¹⁸ F-FDG, including placental crossover ¹⁶						
	Early	3-Month	6-Month	9-Month		
Dose from ¹⁸ F-FDG (mGy/ mCi)	0.81	0.81	0.63	0.63		

PET-CT in the Pregnant Patient: Fetal Dosimetry

Total fetal estimate radiation dose from PET-CT is the sum of CT exposure + maternal gamma irradiation + fetal beta+ + fetal gamma irradiation (Figure 1). Dosimetry estimates from clinical cases are presented in Table 2. Depending on the PET and attenuation correction technique applied, fetal age, and maternal body habitus, fetal estimated doses of between 1.1 and 21.9 mGy have been reported.



Figure 1 - Fetal radiation estimated exposure from ¹⁸F-FDG PET-CT

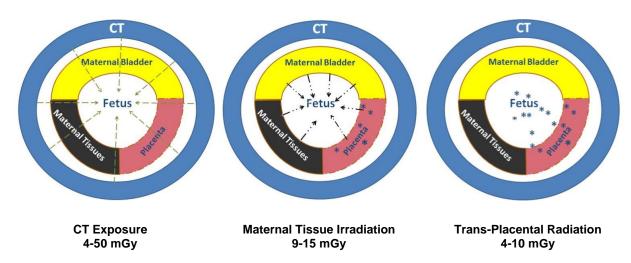


Table 2 - Fetal estimated irradiation from 18F-FDG PET-CT case reports

	Maternal	Placental crossover	СТ	Total
Zanotti-Fregonara et al 2010 ¹⁷	3.0 mGy	8.9 mGy	10 mGy	21.9 mGy
Takalkar et al 2011 ¹⁸	1.1-9.4 mGy (reported combined maternal plus placental crossover)		Negligible (⁶⁸ Ge rod) ¹⁹	1.1-9.4 mGy
Hsieh et al 2012 ²⁰	6.3 mGy (reported combined maternal plus placental crossover)		3.6 mGy	9.9 mGy

PET-CT in the Pregnant Patient: Recommendations

- Establish presence and gestational age of pregnancy
- Establish the indication for medical imaging
- Consider non-ionizing radiation alternative imaging modalities, specifically ultrasound and MRI
- Consider feasibility of obstetrical interventions, including termination or early delivery
- If ¹⁸F-FDG PET is indicated during pregnancy, consider where appropriate:
 - o ¹⁸F-FDG dose reduction to 5 mCi
 - Apply PET imaging techniques such as 3D when possible to support reduced ¹⁸F-FDG dosing.
 - o Increased oral hydration or intravenous saline
 - o Furosemide 20 mg IV, 15 min after ¹⁸F-FDG administration
 - o Encourage frequent voiding.
 - Selective urinary bladder catheterization
 - Attenuation correction with a ⁶⁸Ge rod source is most radiation efficient



- o CT attenuation with 120 kVp and a pitch greater than 1
- o Apply tube current modulation with care to exclude low noise settings
- o Apply iterative CT reconstruction wherever possible
- Estimate total fetal dose: proposed interpretation
 - o Recommendations: < 1 mGy (total gestation)
 - o NRC worker limit: < 5 mGy (0.50 mGy/month)
 - Fetal doses of < 50 mGy, no evidence for fetal injury
 - Fetal doses of < 100 mGy, termination not justified
 - o At fetal doses between 100 and 150* mGy, consider individual circumstances
 - Fetal doses > 150* mGy, possible fetal damage, termination should be seriously considered
 [*equivalent exposure > 3 pelvic CT exams
 - o Fetal doses > 500 mGy, fetal risk sufficient to recommend termination

Additional information is available in the <u>ACR Practice Guideline for Imaging Pregnant Adolescents and Women with</u> Ionizing Radiation.

References

- Wieseler KM, Bhargava P, Kanal KM, et al. Imaging in pregnant patients: examination appropriateness.
 Radiographics, 2010;30:1215-1229; discussion 1230-1213.
 (http://radiographics.rsna.org/content/30/5/1215.abstract)
- 2. Williams PM, Fletcher S. Health effects of prenatal radiation exposure. *Am Fam Physician*, 2010;82:488-493. (http://www.ncbi.nlm.nih.gov/pubmed/20822083)
- 3. Patel SJ, Reede DL, Katz DS, et al. Imaging the pregnant patient for nonobstetric conditions: algorithms and radiation dose considerations. *Radiographics*, 2007;27:1705-1722. (http://www.ncbi.nlm.nih.gov/pubmed/18025513)
- 4. McCollough CH, Schueler BA, Atwell TD, et al. Radiation exposure and pregnancy: when should we be concerned? *Radiographics*, 2007;27:909–917; discussion, 917–918. (http://radiographics.rsna.org/content/27/4/909.full)
- 5. Goldberg-Stein S, Liu B, Hahn PF, Lee SI. Body CT during pregnancy: utilization trends, examination indications, and fetal radiation doses. *AJR*, 2011; 196:146–151. (http://www.ajronline.org/content/196/1/146.full.pdf)
- 6. Goldberg-Stein SA, Liu B, Hahn PF, Lee SI. Radiation dose management: part 2, estimating fetal radiation risk from CT during pregnancy. *AJR*, 2012; 198:W352–W356. (http://www.ncbi.nlm.nih.gov/pubmed/22451572)
- 7. Ratnapalan S, Bona N, Chandra K, Koran G. Physicians' perceptions of teratogenic risk associated with radiography and CT during early pregnancy. *AJR*, 2004;182:1107–1109. (http://www.ajronline.org/content/182/5/1107.full.pdf)
- 8. Bentur Y. Ionizing and nonionizing radiation in pregnancy. In: Koren G, ed. Maternal–fetal toxicology: a clinician's guide, 3rd ed. New York, NY: *Marcel Dekker*, 2001:603–651.
- Angel E, Wellnitz CV, Goodsitt MM, et al. Radiation dose to the fetus for pregnant patients undergoing multidetector CT imaging: Monte Carlo simulations estimating fetal dose for a range of gestational age and patient size. *Radiology*, 2008; 249:220–227. (http://radiology.rsna.org/content/249/1/220.full.pdf+html)



- Lazarus E, DeBenedectis C, North D, Spencer PK, Mayo-Smith WW. Utilization of imaging in pregnant patients: 10-year review of 5270 examinations in 3285 patients—1997–2006. *Radiology*, 2009; 251:517–524. (http://radiology.rsna.org/content/251/2/517.long)
- ACOG Committee on Obstetric Practice. ACOG Committee opinion number 299, September 2004 (replaces no. 158, September 1995): guidelines for diagnostic imaging during pregnancy. *Obstet Gynecol*, 2004; 104:647–651.
 (http://www.acog.org/~/media/Committee%20Opinions/Committee%20on%20Obstetric%20Practice/co299.pdf?dmc=1&ts=20120706T1752458302)
- 12. Webb JA, Thomsen H, Morcos S; The use of iodinated and gadolinium contrast media during pregnancy and lactation. *Eur Radiol*, 2005; 15: 1234–1240. (http://www.ncbi.nlm.nih.gov/pubmed/15609057)
- Bourjeily G, Chalhoub M, Phornphutkul C, Alleyne T, Woodfield C, Chen K. Neonatal thyroid function: effect of a single exposure to iodinated contrast medium in utero. *Radiology*, 2010; 256: 744–750.
 (http://radiology.rsna.org/content/256/3/744.long)
- 14. American College of Radiology (ACR) Website. ACR manual on contrast media, version 7. http://www.acr.org/Quality-Safety/Resources/Contrast-Manual. Accessed 4/11/2012.
- 15. Russell JR, Stabin MG, Sparks RB. Placental transfer of radiopharmaceuticals and dosimetry in pregnancy. *Health Phys*, 1997;73:747–755. (http://www.ncbi.nlm.nih.gov/pubmed/9378650)
- 16. Stabin MG. Proposed addendum to previously published fetal dose estimate tables for 18F-FDG. *J Nucl Med*, 2004;45:634–635. (http://jnm.snmjournals.org/content/45/4/634.full)
- 17. Zanotti-Fregonara P, Jan S, Taieb D, et al. Absorbed 18F-FDG dose to the fetus during early pregnancy. *J Nucl Med*, 2010;51:803-805. (http://jnm.snmjournals.org/content/51/5/803.full.pdf)
- 18. Takalkar AM, Khandelwal A, Lokitz S, et al. 18F-FDG PET in pregnancy and fetal radiation dose estimates. *J Nucl Med*, 2011;52:1035–1040. (http://jnm.snmjournals.org/content/52/7/1035.full.pdf)
- Wu TH, Huang YH, Lee JJ, et al. Radiation exposure during transmission measurements: comparison between CT- and germanium-based techniques with a current PET scanner. *Eur J Nucl Med Mol Imaging*, 2004;31:38–43. (http://www.ncbi.nlm.nih.gov/pubmed/14534833)
- 20. Hsieh TC, MD, Wu YC, Sun SS, et al. FDG PET-CT of a Late-term Pregnant Woman With Breast Cancer. *Clin Nucl Med*, 2012;37: 489–491. (http://www.ncbi.nlm.nih.gov/pubmed/22475901)