MIRD Dose Estimate Report No. 14: Radiation Absorbed Dose from Technetium-99m-Labeled Red Blood Cells

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The estimated absorbed doses from an intravenous administration of technetium-99m-labeled red blood cells are given in Table 1. The data and assumptions used in these calculations are presented as follows.

RADIOPHARMACEUTICAL

Red blood cells (RBCs) may be labeled with technetium-99m (^{99m}Tc) by an in vivo technique, a modified in vivo technique, or an in vitro technique. To label the RBCs in vivo, an intravenous (i.v.) injection of 0.1 mg/kg of stannous pyrophosphate containing 10–20 μ g Sn⁺⁺ is administered to the patient, followed 20–30 min later by the i.v. administration of ^{99m}Tc as sodium pertechnetate (1). A modified technique that results in somewhat higher labeling yields is performed by administering the stannous pyrophosphate intravenously, waiting 20 min, withdrawing 30 ml of blood into a syringe containing heparin and pertechnetate, incubating at room temperature for ~10 min, and then reinjecting the labeled blood intravenously (2).

In vitro labeling is accomplished by the following method. Four ml of blood is withdrawn into a heparinized syringe and transferred to an evacuated vial containing lyophilized stannous citrate. After a 5-min incubation, either 6 ml physiologic saline or 1 ml of 4.4% EDTA solution is added, and the vial is inverted and centrifuged. One to 1.5 ml of red cells is removed and added to a [^{99m}Tc]pertechnetate solution. After another 5-min incubation, the labeled cells are administered intravenously (3).

NUCLEAR DATA

Technetium-99m decays to ⁹⁹Tc by isomeric transition with a half-life of 6.02 hr. Technetium-99 undergoes beta-minus decay with a half-life of 2.13×10^5 yr. The very small contribution of ⁹⁹Tc to the radiation absorbed dose has been ignored in these estimates. The nuclear data for these isotopes are given in Table 2.

 TABLE 1

 Estimated Absorbed Doses from an Intravenous

 Administration of ^{99m}Tc-Bed Blood Cells

	2.4 hr voiding schedule		4.8 hi sch	r voiding nedule			
		In vitro	labeling				
Target	rad/mCi	mGy/MBq	rad/mCi	mGy/MBq			
Heart wall	0.054	0.015	0.054	0.015			
Bladder wall	0.051	0.014	0.087	0.024			
Spleen	0.041	0.011	0.041	0.011			
Lungs	0.041	0.011	0.041	0.011			
Blood	0.035	0.0095	0.036	0.0097			
Liver	0.026	0.0070	0.026	0.0070			
Kidneys	0.025	0.0068	0.025	0.0068			
Red marrow	0.019	0.0051	0.019	0.0051			
Thyroid	0.018	0.0049	0.018	0.0049			
Ovaries	0.017	0.0046	0.018	0.0049			
Testes	0.0071	0.0019	0.0081	0.0022			
Total body	0.015	0.0041	0.015	0.0041			
	In vivo labeling						
Heart wall	0.057	0.015	0.057	0.015			
Bladder wall	0.038	0.010	0.061	0.017			
Spleen	0.043	0.012	0.043	0.012			
Lungs	0.043	0.012	0.043	0.012			
Blood	0.037	0.010	0.038	0.010			
Liver	0.028	0.0076	0.028	0.0076			
Kidneys	0.027	0.0073	0.027	0.0073			
Red marrow	0.020	0.0054	0.020	0.0054			
Thyroid	0.019	0.0051	0.019	0.0051			
Ovaries	0.018	0.0049	0.019	0.0051			
Testes	0.0076	0.0021	0.0082	0.0022			
Total body	0.016	0.0043	0.016	0.0043			

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TABLE 2	
Nuclear Data	

 Radionuclide	<u></u>		^{99m} Tc		⁹⁹ Tc			
Physical half-life			6.02 h		2.13 × 10⁵ y			
Decay constant			0.1151 h ^{−1}		$3.25 \times 10^{-6} \text{ y}^{-1}$			
Mode of decay			I.T.		β-			
Principal Radiations	E,	n,	Δ_{i}		Δ_{i}			
						(Gy kg/MBq		
	(keV)		(rad g/µCi h)	(Gy kg/MBq s)	(rad g/µCi h)	s)		
Photon	18–21	0.079	0.0029	2.18 × 10 ⁻¹⁰				
	140.5	0.89	0.266	2.00 × 10 ⁻⁸				
Nonpenetrating			0.0332	2.49 × 10 ⁻⁹	0.216	1.62 × 10 ^{−8}		

E_i is energy per photon.

n, is mean number of particles or photons per nuclear transition.

 $\Delta_{\!i}$ is mean energy emitted per nuclear transition.

Nonpenetrating radiation from 99mTc includes conversion and Auger electrons ranging in energy from 1.6 keV to 140 keV. Nonpenetrating radiation from 99Tc includes beta-minus emissions with a maximum energy of 294 keV and an average energy of 101.3 keV. Only photons whose mean number per transition is 0.01 or greater are included. See references (6, 7) for sources of nuclear data.

Note: Complete decay of 1 unit of activity of ^{99m}Tc produces 3.2×10^{-9} units of ⁹⁹Tc.

BIOLOGIC DATA

These dose estimates are based on data obtained from nine normal subjects studied with the in vivo and in vitro labeling techniques at Brookhaven National Laboratory (4). Biologic parameters and residence times listed in Table 3 were calculated by curve fitting of the urine and blood data from these nine subjects. Corroborative data were obtained from subjects studied at the National Institutes of Health and at Albert Einstein College of Medicine, Bronx, NY.

With the in vivo technique, a variable percentage

(from 60%-90%) of the administered activity labels the circulating RBCs after the i.v. injection of [99mTc]pertechnetate. The dose estimates in this report use an observed in vivo labeling efficiency of 84% (4). The maximum level of radioactivity associated with the RBCs is attained 30-60 min after the 99mTc administration. The bulk of the activity not attached to the RBCs will be excreted, probably as a nonpertechnetate form of technetium after distribution throughout the extracellular space. This is considered to be in the "remainder of the body" prior to excretion. A small and variable fraction of the activity may remain as pertechnetate. In

Biologic Parameters							
Organ	α _{h1}	λ _{h1} h ⁻¹	α2	λ _{n2} h ⁻¹	α3	λ _{n3} h ⁻¹	τ _n h
		In	vitro labeling				
Blood		_	0.380	0.694	0.629	0.0339	5.97
Bladder contents							
2.4 hr void	_	_	—	—			0.23
4.8 hr void	_	-		—	_		0.45
Total body	_	_	0.273	0.186	0.727	0.00481	7.05
Remainder of the body		—	_	_	_	-	1.08
		In	vivo labeling				
Blood	-0.174	3.13	0.375	0.211	0.525	0	6.19
Bladder contents							
2.4 hr void		_			_		0.14
4.8 hr void		—		—			0.28
Total body		_	0.126	0.279	0.874	0.00450	7.69
Remainder of the body					—		1.50

TABLE 3

The r-value is given as the average of the individual patient values and can differ somewhat from the value which would be calculated from the average α and λ .

 TABLE 4

 S Values for Various Source Organs (rad/mCi hr)

	Source Organs					
Target	Blood ^(a)	Bladder contents ^(c)	Remainder of the body ⁽¹⁾			
Heart wall	8.8 × 10 ⁻³	$4.4 imes 10^{-5(d)}$	1.7 × 10 ⁻³			
Bladder wall	2.1 × 10 ^{−3(b)}	1.6 × 10⁻¹	1.8 × 10 ^{−3}			
Spleen	6.5 × 10 ^{−3}	6.6 × 10 ⁻⁴	1.9 × 10 ⁻³			
Lungs	6.6 × 10⁻³	2.4 × 10⁻⁵	1.6 × 10 ⁻³			
Blood	5.5 × 10⁻³	2.1 × 10 ^{−3(e)}	2.0 × 10 ^{−3(g)}			
Liver	4.0 × 10 ^{−3}	1.7 × 10⁻⁴	2.1 × 10 ^{−3}			
Kidneys	3.8 × 10 ^{−3}	2.6 × 10 ⁻⁴	2.1 × 10 ^{−3}			
Red marrow	2.5 × 10 ⁻³	2.2 × 10 ⁻³	2.9 × 10 ^{−3}			
Thyroid	2.7 × 10⁻³	2.1 × 10 ⁻⁶	1.4 × 10 ^{−3}			
Ovaries	2.1 × 10 ⁻³	7.3 × 10 ⁻³	2.4 × 10 ^{−3}			
Testes	6.9 × 10 ^{-₄}	4.7 × 10 ⁻³	1.8 × 10 ⁻³			
Total body	2.0 × 10 ⁻³	1.9 × 10 ⁻³	2.0 × 10 ⁻³			

Note: The values in the above table are in units of rad/mCi hr. To change to rad/ μ Ci hr, multiply by 10⁻³ (e.g., S (heart walk-blood) = 8.8 × 10⁻⁶ rad/ μ Ci hr)

 $^{\rm a}$ S-values for the blood as a source organ were taken from Ref. 5.

^b Taken to be the same as the S-value for blood as the source organ to the uterus as the target.

 $^{\rm c}$ Except as indicated, S-values were taken from MIRD Pamphlet 11 (Ref. 8).

^d Derived from Mird Pamphlet 13 (Ref. 9).

^e Taken to be the same as the S-value for blood as the source organ and the uterus as the target organ using the reciprocity relationship.

¹ Remainder of the body refers to the total body minus the blood and bladder contents. The remainder of the body S-value for each target organ, except the blood, is obtained by multiplying the total body to organ S-value by the mass ratio of total body to remainder of the body and then subtracting the blood to organ and bladder contents to organ S-values, each multiplied by their respective source organ to remainder of the body mass ratios (Ref. 10).

⁹ Taken to be the same as the S-value for blood as the source organ and the total body as the target organ using the reciprocity relationship.

addition, technetium is gradually eluted from the labeled RBCs and excreted in the urine. A total of $\sim 21\%$ of the injected activity (unbound as well as eluted from red cells) is excreted over a 24-hr period.

Labeling efficiency with the in vitro method is generally ~97%-98% (3). Immediately after administration of the labeled blood, there is some elution (~5%) of ^{99m}Tc. Peak blood levels are obtained immediately after administration, whereas peak levels following administration of activity with the in vivo method are significantly delayed. Peak blood levels with the in vitro labeling technique are higher $(24 \pm 16\%)$ than those with the in vivo technique. For the in vitro labeling method, blood levels remain higher for up to 8 hr, after which they are lower in comparison to the in vivo method because of a higher excretion rate (4).

ABSORBED DOSE ESTIMATES

Absorbed dose calculations are based on the biologic parameters in Table 3. Source organs are blood, bladder contents, and "remainder of the body." "Remainder of the body" distribution is assumed for that fraction which is not in the blood pool and which has not been excreted. The S values for source organs are given in Table 4.

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