Partial-body dose estimation

Feature present since ADCI version 1.12.

Manuscript presenting the partial-body algorithm in ADCI: Direct link to pdf

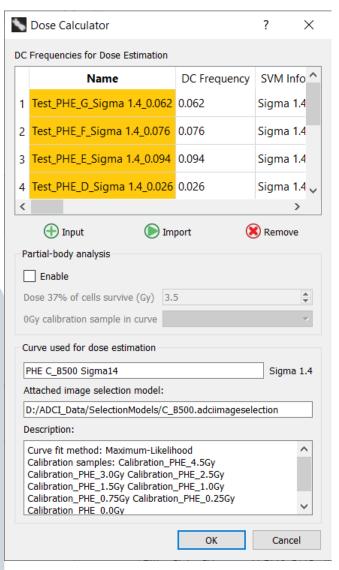
Enable partial-body dose estimation

Partial-body dose estimation can be performed on processed test samples. It can be enabled by first executing the dose estimation wizard (recommended method) or directly through the dose calculator. Whether the dose estimation wizard was executed first or not, clicking the checkbox labeled "enable" in the Dose Calculator dialog will instruct ADCI to perform partial-body analysis in addition to standard whole-body dose estimation.

> Partial-body dose estimation requires a calibration curve with both an associated SVM sigma value and image selection model. These conditions will almost always be met if the calibration curve was generated using the curve calibration wizard. In the event either contidion is not met, the "enable" checkbox will be disabled and a tooltip describing the unmet condition will be displayed if the mouse is hovered over the disabled checkbox.



Test samples can be imported in the same manner as standard whole-body dose estimation. Consult documentation related to the dose estimation wizard (recommended method) and how to automatically import test samples to the Dose Calculator. Manual entery of dicentric frequencies is not compatible with partial-body dose estimation as the distribution of DCs across all images in a sample is required.



Enable partial-body dose estimation (contd)

After clicking the "Enable" checkbox, the dose at which 37% of irradiated cells survive must be provided. It is pre-populated based on the value specified in the "Partial-body Preferences" dialog described at the end of this page. The value can be adjusted here either by typing in the desired value or by using the mouse wheel while hovering the mouse above the "Dose 37% of cells survive" spinbox.

The OGy calibration sample used to generate the calibration curve must be selected from the "OGy calibration sample in curve" dropdown. All samples currently in the main GUI sample list are present in the dropdown.

It is important to ensure the OGy calibration sample used to generate the curve is selected in the "OGy calibration sample in curve" dropdown. ADCI attempts to prepopulate the dropdown with the appropriate sample, however it does this by attempting to parse sample filenames and may not always pre-populate the dropdown correctly. Additionally, the OGy calibration sample must first be present in the main GUI sample list in order to appear in the dropdown.

Once all desired test samples are present in the Dose Calculator, a desired value for "Dose 37% of cells survive" is specified, and the appropriate OGy calibration sample is selected in the "OGy calibration sample in curve" dropdown click the "OK" button to begin the analysis. After a short delay, a progress bar will display the progress of the dose estimation.

0	Dose Calculator		ΥΥΥΥΥ				
DC Frequencies for Dose Estimation							
	Name	DC Frequency	SVM Info				
1	Test_PHE_G_Sigma 1.4_0.062	2 0.062	Sigma 1.4				
2	Test_PHE_F_Sigma 1.4_0.076	0.076	Sigma 1.4				
3	Test_PHE_E_Sigma 1.4_0.094	0.094	Sigma 1.4				
4	Test_PHE_D_Sigma 1.4_0.020	<mark>5</mark> 0.026	Sigma 1.4 🗸				
<		>					
Partial-body analysis Enable Dasa 27% of colls suprise (Cu) 2.5							
Dose 37% of cells survive (Gy) 3.5 0Gy calibration sample in curve Calibration_PHE_0.0Gy							
Curve used for dose estimation Curve used for dose estimation PHE C_B500 Sigma14 Attached image selection model: D:/ADCI_Data/SelectionModels/ Test_PHE_A Calibration_PHE_0.0Gy Test_PHE_D Test_PHE_D Test_PHE_B Test_PHE_A							
D	escription:						
Curve fit method: Maximum-Likelihood Calibration samples: Calibration_PHE_4.5Gy Calibration_PHE_3.0Gy Calibration_PHE_2.5Gy Calibration_PHE_1.5Gy Calibration_PHE_1.0Gy Calibration_PHE_0.75Gy Calibration_PHE_0.25Gy Calibration PHE 0.0Gy							
		ОК	Cancel				

Interpret partial-body dose estimation results

Please note the content of the plot area in the main GUI reflects the results of standard whole-body

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dose estimation and not partial-body dose estimation. Whether partial-body analysis is enabled or not, the plot will always reflect whole-body results.

Partial-body dose estimation results are appended to standard whole-body results in the console area within the main GUI. Example console output is shown below. These output fields also appear in dose estimation reports if partial-body analysis was performed and are available as user-selected fields for provider (csv) dose estimation reports.

ainty PB Estimated Dose	PB Dose LCL	PB Dose UCL	PB Cells Irradiated	PB Cells Irradiated LCL	PB Cells Irradiated UCL	Applied Image Selection
+ Curve 4.20Gy	3.00Gy	5.15Gy	55.84%	35.80%	80.92%	Images are selected by
+ Curve 3.15Gy	1.60Gy	4.15Gy	57.46%	30.25%	*100.00%	Images are selected by
+ Curve 1.75Gy	*0.00Gy	2.85Gy	78.14%	31.60%	*100.00%	Images are selected by

• PB Estimated Dose

Estimated dose (Gy) of the irradiated fraction.

• PB Estimated Dose LCL

Estimated dose (Gy) of the irradiated fraction lower confidence limit. The mean dicentric yield of the irradiated fraction is calculated as part of the Contaminated poisson method. The mean dicentric yield minus one standard deviation is compared to the curve to arrive at this value.

• PB Estimated Dose UCL

Estimated dose (Gy) of the irradiated fraction upper confidence limit. The mean dicentric yield of the irradiated fraction is calculated as part of the Contaminated poisson method. The mean dicentric yield plus one standard deviation is compared to the curve to arrive at this value.

PB Cells Irradiated Estimated percentage of total cells exposed.

 PB Cells Irradiated LCL Calculated using PB Estimated Dose UCL in place of PB Estimated Dose
 PB Cells Irradiated UCL

Calculated using PB Estimated Dose LCL in place of PB Estimated Dose

In the event the Contaminated poisson method cannot be performed, all fields listed above will read "Err". At least one image in a test sample must contain >1 DC in order for the method to be performed. Additionally, because of this requirement the Contaminated poisson method cannot be performed on a manually entered DC frequency.

Above this table, and additional supplementary table labelled "Extended partial-body data" lists values used in the Contaminated poisson method. These fields are: DC Yield in Irradiated Fraction, Fraction of Surviving Cells Reaching Metaphase, Fraction of Irradiated Cells, Dose of Irradiated Fraction, Fraction, Fraction of Cells Exposed.

Access parameters dialog

Access the "Partial-body Parameters" dialog through the menu bar location at the top of the main gui. Click "Settings" and then select "Partial-Body Preferences".

	🔪 Adci								
	Samples	Curves	ADCI Process	Queue	Wizards	Reports	Settings	Help	
						Change FP Filters		Ì	
	Sample	Samples						Statistics Options	
	Ident	dentity Size Processed	Size Processed			ADCI Preferences		t	
						Qual	ity Control Thresholds	L	
					Image Selection Optimization Settings		L		
					GPU Settings		I		
						Partia	al-Body Preferences		

Brief discussion of algorithm

A more detailed explanation of the partial-body algorithm utilized by ADCI can be found in our

manuscript published in the International Journal of Radiation Biology (2020): Direct link to pdf

Although FP DCs in ADCI are minimized by morphological image filtering, they are not completely eliminated. The impact of remaining FP DCs on dose estimates of partially irradiated samples was not predictable and affected the accuracy of some estimates, especially at low-dose exposures. Here, we correct the overall DC frequencies by removing the estimated residual FPs after morphological filtering by assuming nearly all DCs detected in the 0Gy calibration sample are FPs. ADCI generates a user-defined number (default: 500) of randomly selected subsets of samples, each subset containing half the number of images present in the original 0Gy calibration sample, or a user-defined number (default: 500) of images, whichever is greater. ADCI then determines the DC frequency of each subset (after application of an image selection model) and calculates the standard deviation. A calibration curve adjusted for FPs is generated by reducing the observed 0Gy calibration sample DC frequency by two SD. DCs are then removed from all other calibration samples based on the residual DC count in the 0Gy sample. The 0Gy calibration sample DC frequency reduced by 2 SD is a conservative estimate of the minimum DC frequency which could reasonably be found in the sample. By removing this reduced number of FPs in test samples, we ensure that DC counts in test samples after FP adjustment are non-negative.

Description of settings

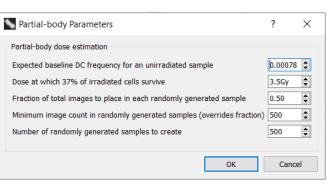
• Expected baseline DC frequency for an unirradiated sample

Baseline DC frequency in unirradiated cells varies between studies, reported to range from 0.00009 to 0.00299¹⁾. Baseline DC frequencies may vary between laboratories. Keep in mind this value does not reflect the DC frequency reported by ADCI for an unirradiated sample, instead it is the DC frequency for an unirradiated sample scored by an expert. We have selected a default baseline DC frequency of 0.00078 based on Lloyd et al. 1980²⁾.

• Dose at which 37% of irradiated cells survive

Corresponds to the term D_0 in the Contaminated poisson method as described within the IAEA manual in section 9.7.4.3³⁾ and varies based on the type of radiation to which a individual was exposed. The default value for this parameter is 3.5Gy.

• Fraction of total images to place in each randomly generated sample Random subsets of images from an unirradiated sample are derived in order to determine an expected minimum number of DCs. A fraction of 0.5 means each randomly selected subset contains 50% of the images found in the original



unirradiated sample. The default value for this parameter is 0.5.

Minimum image count in randomly generated samples (overrides fraction)

A minimum image count for each randomly generated subset. For example, if a 0Gy calibration sample contains 1000 images and a fraction of 0.4 is applied, 400 images will be present in each subset. Then, if the minimum image count specified here is higher than 400, the minimum image count will override the fraction and each randomly generated subset will contain the minimum image count. Additionally, if the minimum image count is higher than the number of images in the 0Gy calibration sample, dose estimation will be aborted. The default value for this parameter is 500.

• Number of randomly generated samples to create

Count of randomly generated subsets to create. A higher number will increase dose estimation consistency but will require more computation time. The default value for this parameter is 500.

1)

Romm H, Oestreicher U, Kulka U. Cytogenetic damage analysed by the dicentric assay. Annali Dell'istituto Superiore di Sanita. 2009 ;45(3):251-259.

2)

Lloyd DC, Purrott RJ, Reeder EJ. The incidence of unstable chromosome aberrations in peripheral blood lymphocytes from unirradiated and occupationally exposed people. Mutat Res. 1980;72(3):523-532. doi:10.1016/0027-5107(80)90123-2

3)

https://www-pub.iaea.org/MTCD/Publications/PDF/EPR-Biodosimetry%202011_web.pdf