SUPPLEMENTAL TABLE 1a.

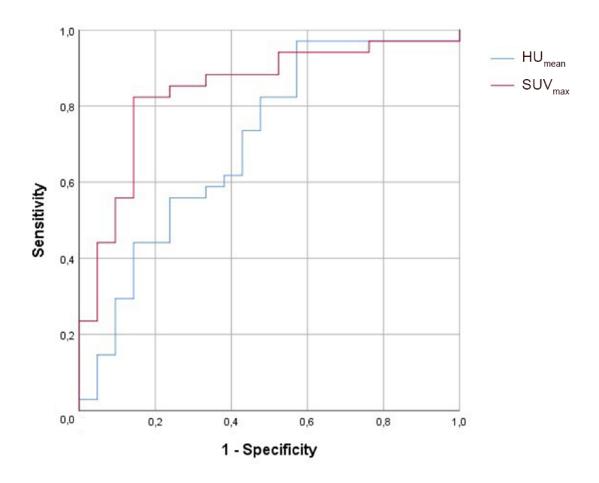
Univariate logistic regression analysis of biopsy outcome and significant continuous variables. N=55 biopsies.

Variable	р	Odds ratio	95% CI
SUV _{mean}	0.016	4.509	1.324-15,355
SUV _{max}	0.008	6.889	1.646-28.834
HU _{mean}	0.006	0.996	0.993-0.999
HU _{max}	0.037	0.998	0.995-1.000

SUPPLEMENTAL TABLE 1b.

Multivariate logistic regression analysis of biopsy outcome and significant continuous variables. N=55 biopsies.

Variable	р	Odds ratio	95% CI
SUV _{max}	0.003	11.737	2.258-60.996
HU _{mean}	0.003	0.995	0.992-0.998



SUPPLEMENTAL FIGURE 1.

ROC curves for biopsy outcome by HU_{mean} and SUV_{max} at biopsy site. N=55 biopsies.



SUPPLEMENTAL FIGURE 2.

Mutational landscape of ⁶⁸Ga-PSMA guided bone biopsies identified by wholegenome sequencing.

- A) Estimated tumor cell percentages (*in silico*) obtained from the whole-genome sequencing data. Bars are color-coded based on tumor cell percentage.
- B) Number of genomic mutations per Mbp (TMB) of SNV, InDels, and MNV categories. All genome-wide somatic mutations were taken into consideration (square root scale). C) Absolute number of unique structural variants per sample. Cumulative frequency of inversions, tandem duplication, deletions, insertions, and translocations.
- D) Relative frequency per structural variant category. Tandem Duplications and Deletions are subdivided into >100 kbp and <100 kbp categories. This track shows whether enrichment for a particular category of (somatic) structural variant can be detected, which in turn, can be indicative for a specific mutational aberration.
- E) Relative genome-wide ploidy status, ranging from 0 to ≥7 copies. This track shows the relative percentage of the entire genome which is (partially) deleted (ploidy < 2/diploid) or amplified (> 2/diploid).
- F) Relative contribution to mutational signatures (COSMIC) summarized per proposed etiology. This track displays the proposed etiology of each SNV based on their mutational contexts. Signatures with < 5% relative contribution in all samples were summarized in the "Filtered (<5%)" category.
- G) Relative frequency of somatic Single Nucleotide Variants (SNV) categories.
- H) Relative frequency of somatic Doublet Base Substitution (DBS) categories.
- I) Oncoplot showing the somatic mutations per samples for a selection of potential driver genes, as detected by dN/dS and/or GISTIC2 analysis and/or manual selection based on

a list of pan-cancer (CPCT-02) driver genes¹. Shallow amplifications and deletions are only shown if also accompanied by an additional coding mutation. The right-hand bar-plot depicts the relative frequency of mutational categories (coding mutations, structural variants and deep amplifications/deletions) per gene. Per gene, the inclusion criteria are shown; dN/dS ($q \le 0.1$; *), enriched GISTIC2 focal peak ($q \le 0.1$; †) or if empty, based on the pan-cancer (CPCT-02) driver list.

- J) Presence of chromothripsis. Pink color indicates presence of chromothripsis as estimated by ShatterSeek.
- K) Presence of kataegis. Red color indicates presence of one or more regions showing kataegis.
- L) Presence of a fusion with a member of the ETS transcription factor family. Green color indicates a possible fusion.