

TAKE-HOME MESSAGES

- ❑ **SPECT-like DVHs** largely collapsed toward **Dmean**, offering little added information beyond mean-dose reporting.
- ❑ **PET-like DVHs** remained imperfect, but may preserve **residual heterogeneity information** under selected conditions relevant to **Y-90 TARE personalization**.

INTRODUCTION

- ❑ Dose Volume Histogram (DVH) metrics such as D90 and D10 are increasingly reported in Y-90 TARE dosimetry.
- ❑ Unlike external beam radiation therapy, Y-90 DVHs are derived from **limited-resolution** PET or SPECT images.
- ❑ **When heterogeneity is not spatially resolved, D90 and D10 may become largely predictable from Dmean.** In that setting, DVH metrics may be calculable but not independently informative.
- ❑ **Aim:** Determine how tumor size, intratumoral heterogeneity, and image resolution affect the reliability of image-based D90 and D10 in Y-90 dosimetry.
- ❑ **Significance:** Define when D90 and D10 are true heterogeneity metrics versus resolution-limited surrogates of Dmean.

MATERIALS AND METHODS

- ❑ **Analytically defined 3D digital phantoms** were generated in Python to model tumor geometry and intratumoral heterogeneity under controlled conditions.
- ❑ Models included **uniform spheres**, **cold regions**, and **hot regions**, providing known ground truth for DVH analysis.
- ❑ **Y-90-like reference dose maps** were generated using a **3-mm Gaussian kernel**; broader FWHM kernels simulated **PET-** and **SPECT-like** image degradation.
- ❑ Dmean, D90, and D10 were extracted from each simulated dose map.
- ❑ **R²** were used to quantify when D90 and D10 became redundant with Dmean.

KEY RESULTS AND SUMMARY

- ❑ DVHs were affected by **lesion size and effective resolution** even in uniform activity spheres.
- ❑ For spheres with cold cores, worsening resolution increased **D90 redundancy with Dmean**.
- ❑ For spheres with hot spots, worsening resolution increased **D10 redundancy with Dmean**.
- ❑ Image-based DVH metrics are most meaningful when the hot or cold regions they describe remain **spatially resolvable**.
- ❑ DVH interpretation in Y-90 dosimetry should account for **tumor size, heterogeneity scale, and effective image resolution**.

RESULTS AND DISCUSSION

- Uniform Activity Spheres: DVH Depends on Lesion Size and Effective Resolution*
 - ❑ **DVH fidelity decreases** as lesion size approaches the effective imaging resolution.
 - ❑ **PET-like resolution (5–10 mm)** largely preserves D90/Dmean trends for larger lesions, whereas **SPECT-like resolution (15–20 mm)** progressively obscures them.
 - ❑ **Loss of monotonicity** between blurred and reference D90/Dmean indicates that the DVH metric has become **resolution-dominated** rather than reflective of the underlying dose distribution.

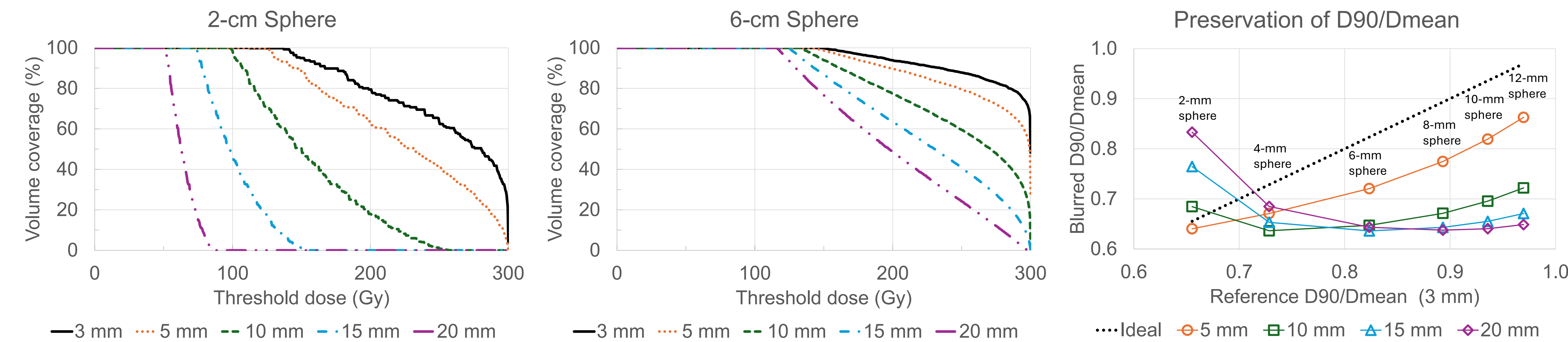


Figure 1. Effect of spatial resolution on DVH metrics in uniform spheres. Left/Center: DVHs for 2- and 6-cm spheres using 3-mm reference, 5–10 mm PET-like, and 15–20 mm SPECT-like Gaussian blurring. Right: Blurred D90/Dmean versus the corresponding 3-mm reference value for sphere diameters of 2–12 cm; the identity line indicates perfect agreement.

2. Spheres w/ Cold Spots: Spatial Blurring Makes D90 Increasingly Redundant with Dmean

- ❑ At the **3-mm reference resolution**, cold-spot geometry produced substantial variation in D90 among lesions with similar Dmean.
- ❑ This variability remained **appreciable at 10-mm PET-like resolution**, but became **minimal at 20-mm SPECT-like resolution**.
- ❑ Thus, worsening spatial resolution progressively converts D90 from a heterogeneity-sensitive metric into a predictable transformation of Dmean.

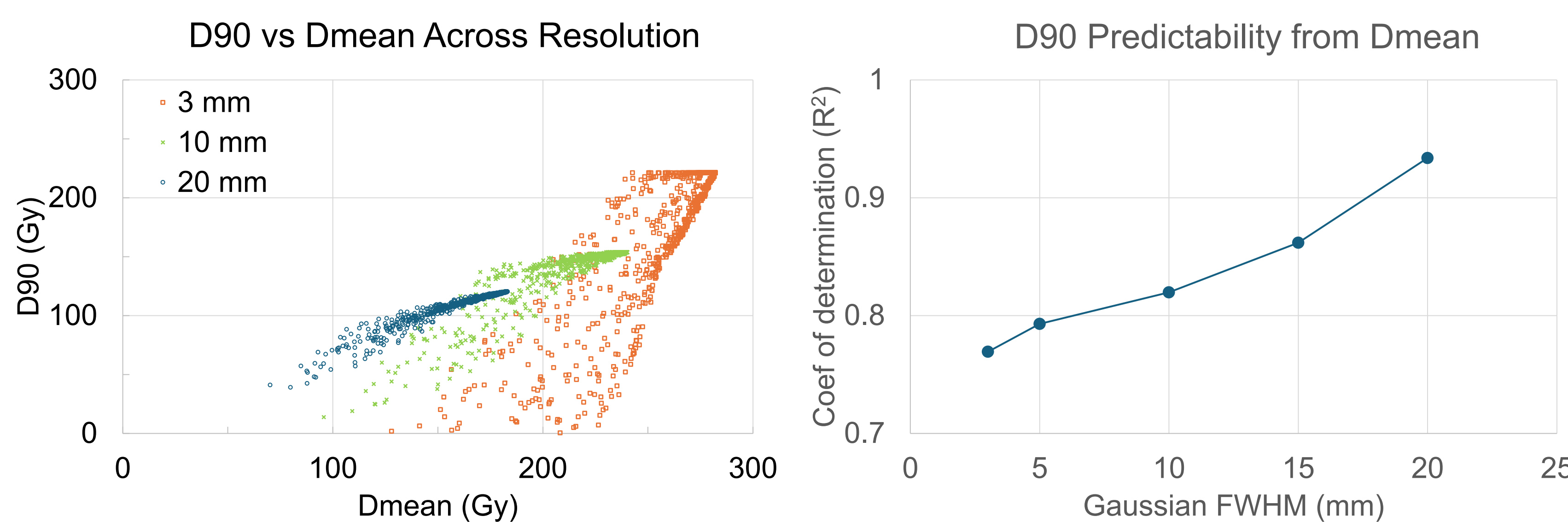


Figure 2. Spatial-resolution dependence of the Dmean–D90 relationship in cold-spot simulations. 1000 5-cm spheres contained a single randomly located cold spot with diameter 4–40 mm and relative uptake 0–1. Left: The broad Dmean–D90 spread at the 3-mm reference resolution narrowed progressively at 10 and 20 mm. Right: R² increased from 0.77 to 0.93, quantifying the increasing predictability of D90 from Dmean.

3. Spheres w/ Hot Spots: Spatial Blurring Makes D10 Increasingly Redundant with Dmean

- ❑ At the **3-mm reference resolution**, hotspot geometry produced substantial variation in **D10** among lesions with similar Dmean, indicating preserved high-dose heterogeneity information.
- ❑ This D10 variability **persisted at 10-mm PET-like resolution**, but was largely **lost at 20-mm SPECT-like resolution**.
- ❑ Thus, worsening spatial resolution progressively converts **D10** from a hotspot-sensitive DVH metric into a predictable transformation of Dmean.

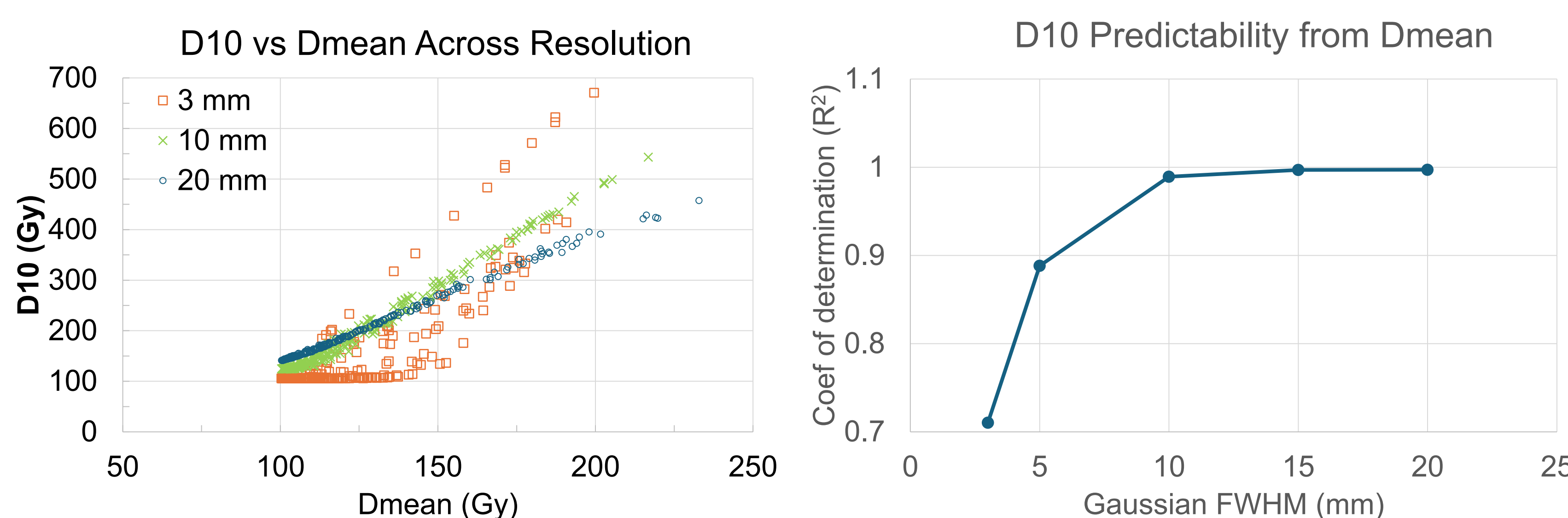


Figure 3. Spatial-resolution dependence of the Dmean–D10 relationship in hot-spot simulations. 300 5-cm spheres contained a single randomly located hotspot with radius 3–12 mm and relative uptake 2–10. Left: The broad Dmean–D10 spread at the 3-mm reference resolution narrowed progressively at 10 and 20 mm. Right: R² increased from 0.71 to ~1.00, quantifying the increasing predictability of D10 from Dmean.