

Development of regime map for steady-state high shear wet twin-screw granulation

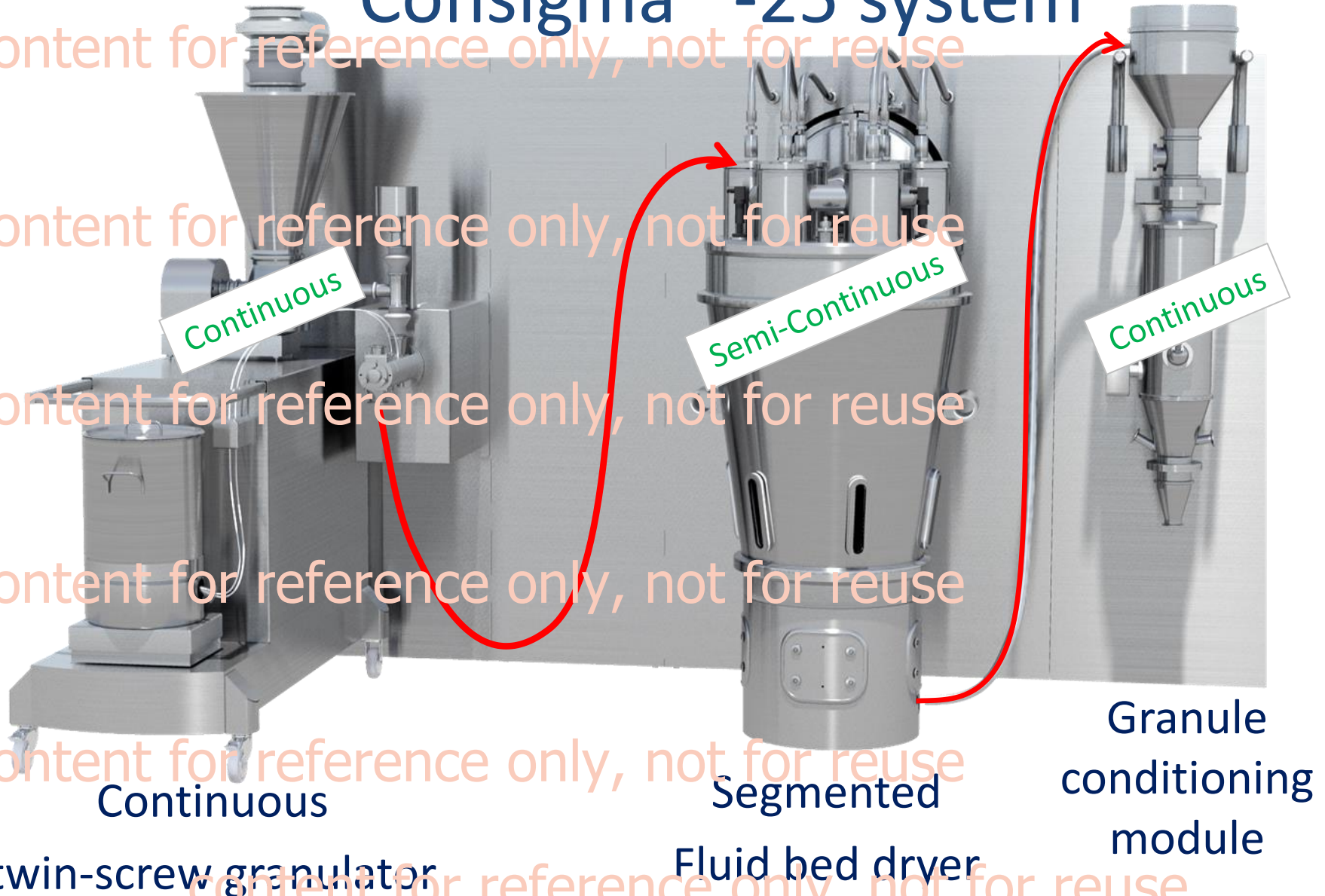
Ashish Kumar

7th International Granulation Workshop



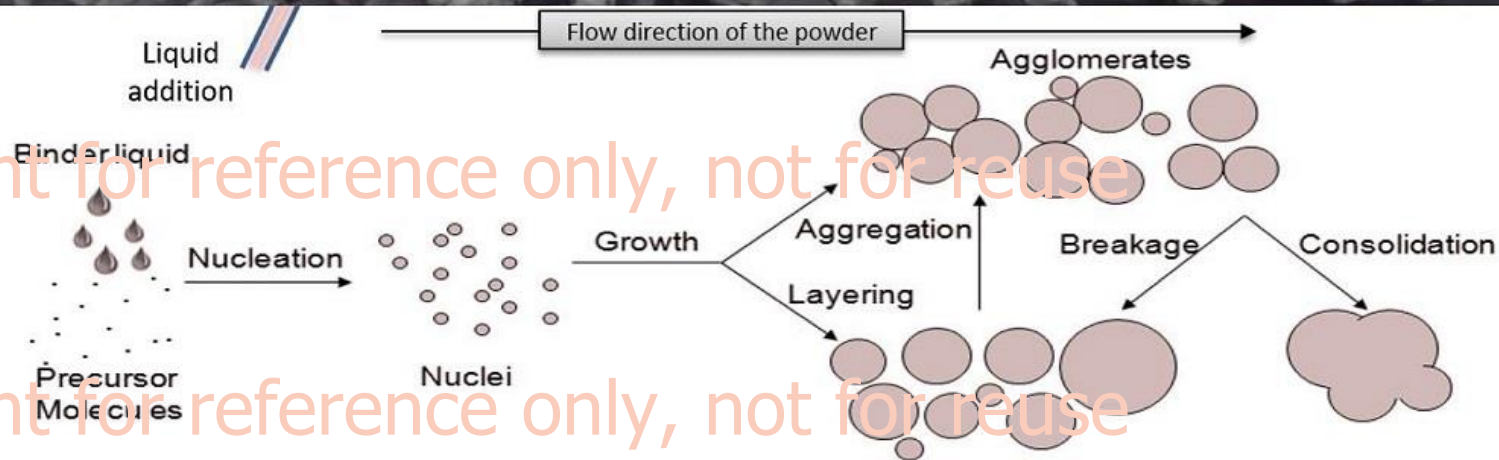
Continuous manufacturing line

Consigma™-25 system



High shear wet granulation in TSG involves

many different process and equipment variables



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Challenges in wet granulation

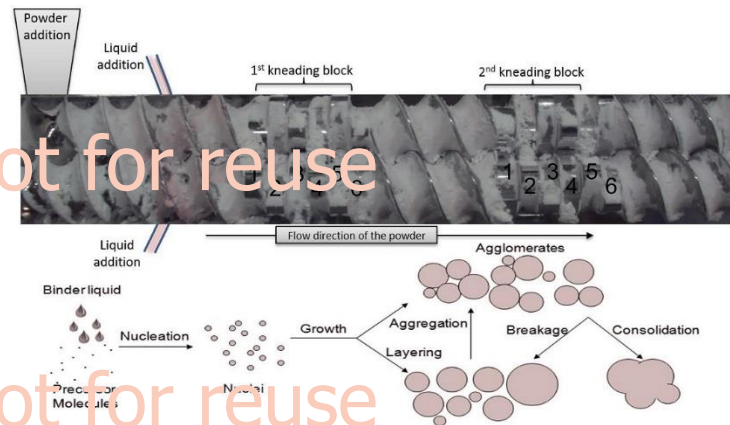
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Understanding the effect of process settings on granulation performance

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Process evolution

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Product quality

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Mostly theoretical approaches tracking granule attributes

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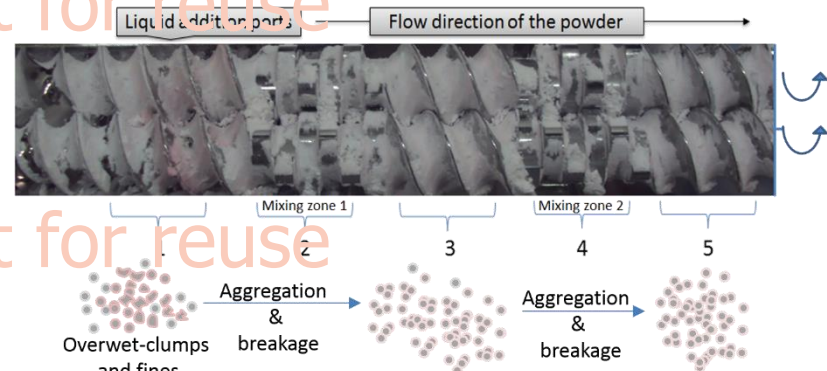
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Challenges in wet granulation

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Understanding the effect of process settings on granulation performance

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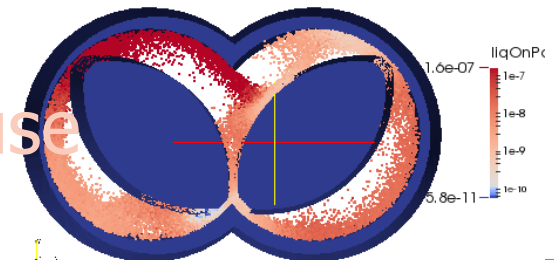
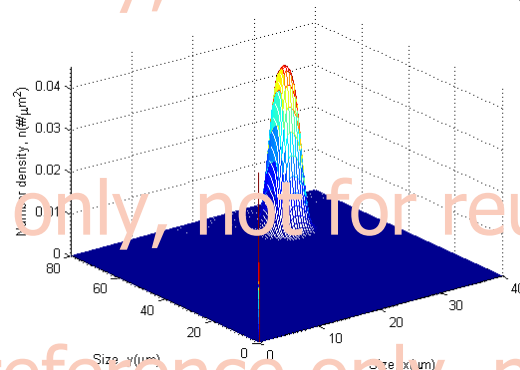
- **Process evolution**

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Mostly theoretical approaches tracking granule attributes

- **Product quality**

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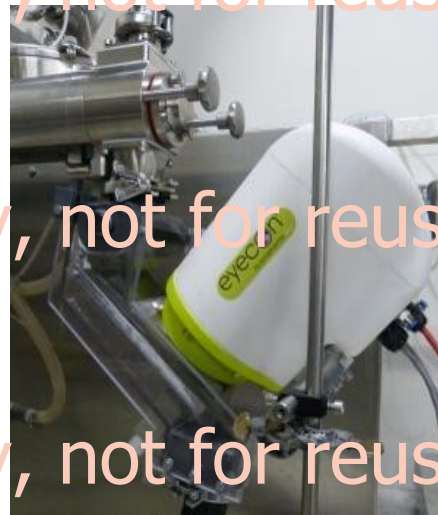
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Challenges in wet granulation

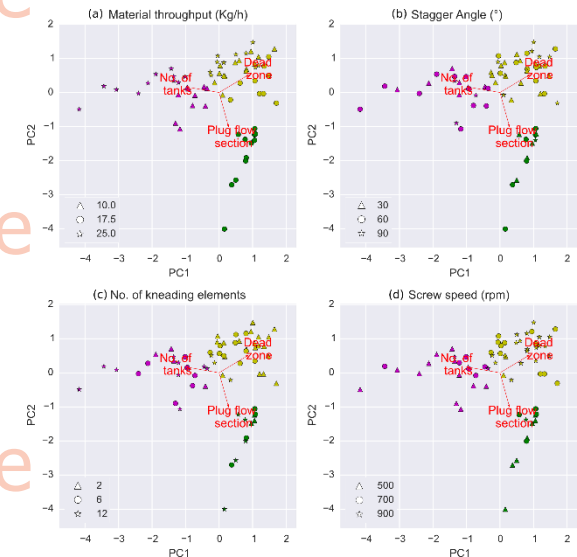
Understanding the effect of process settings on granulation performance

- Process evolution

- Product quality



Experiments



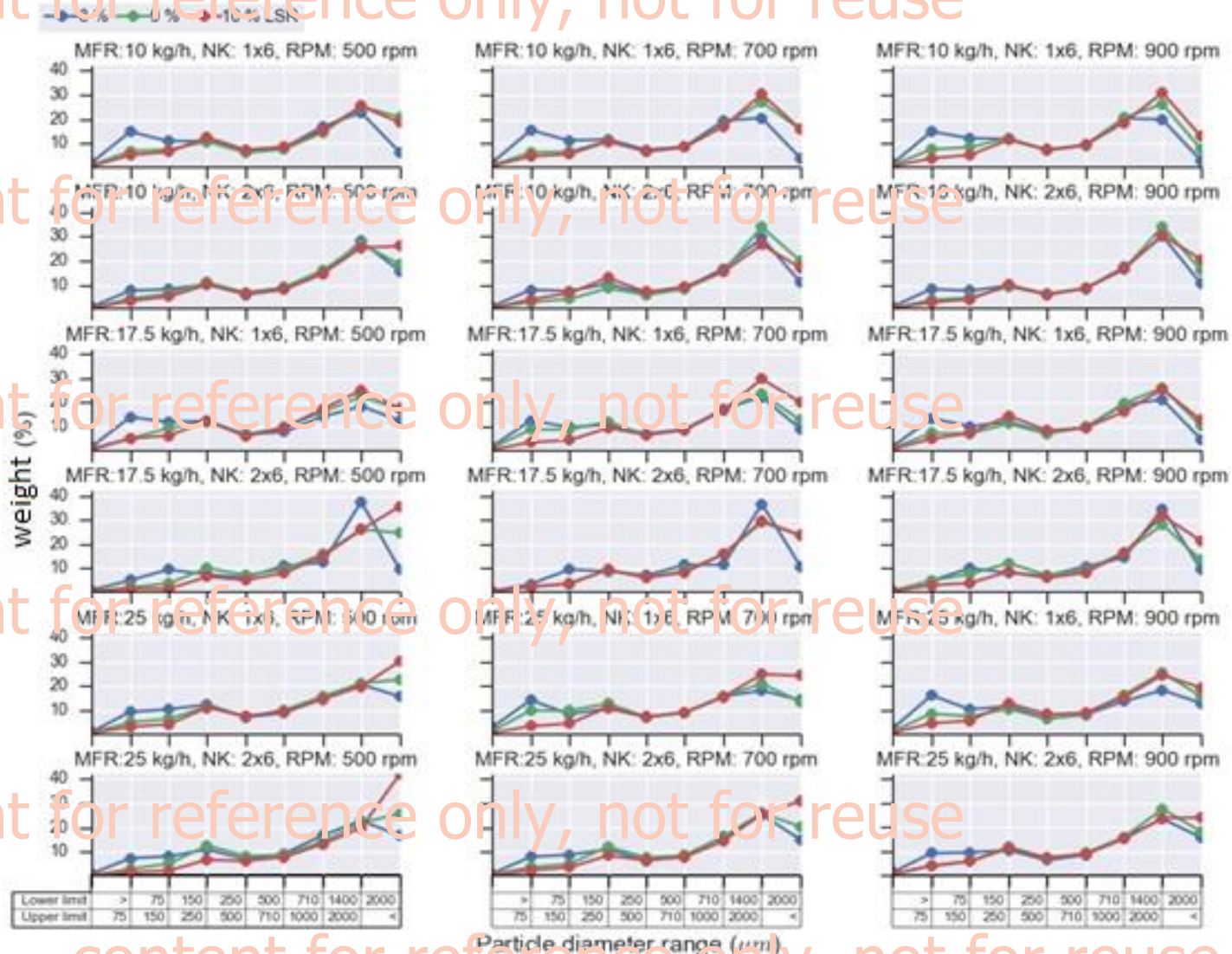
multivariate data analysis

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Traditional approach involves change in

scale dependent process settings on PSD

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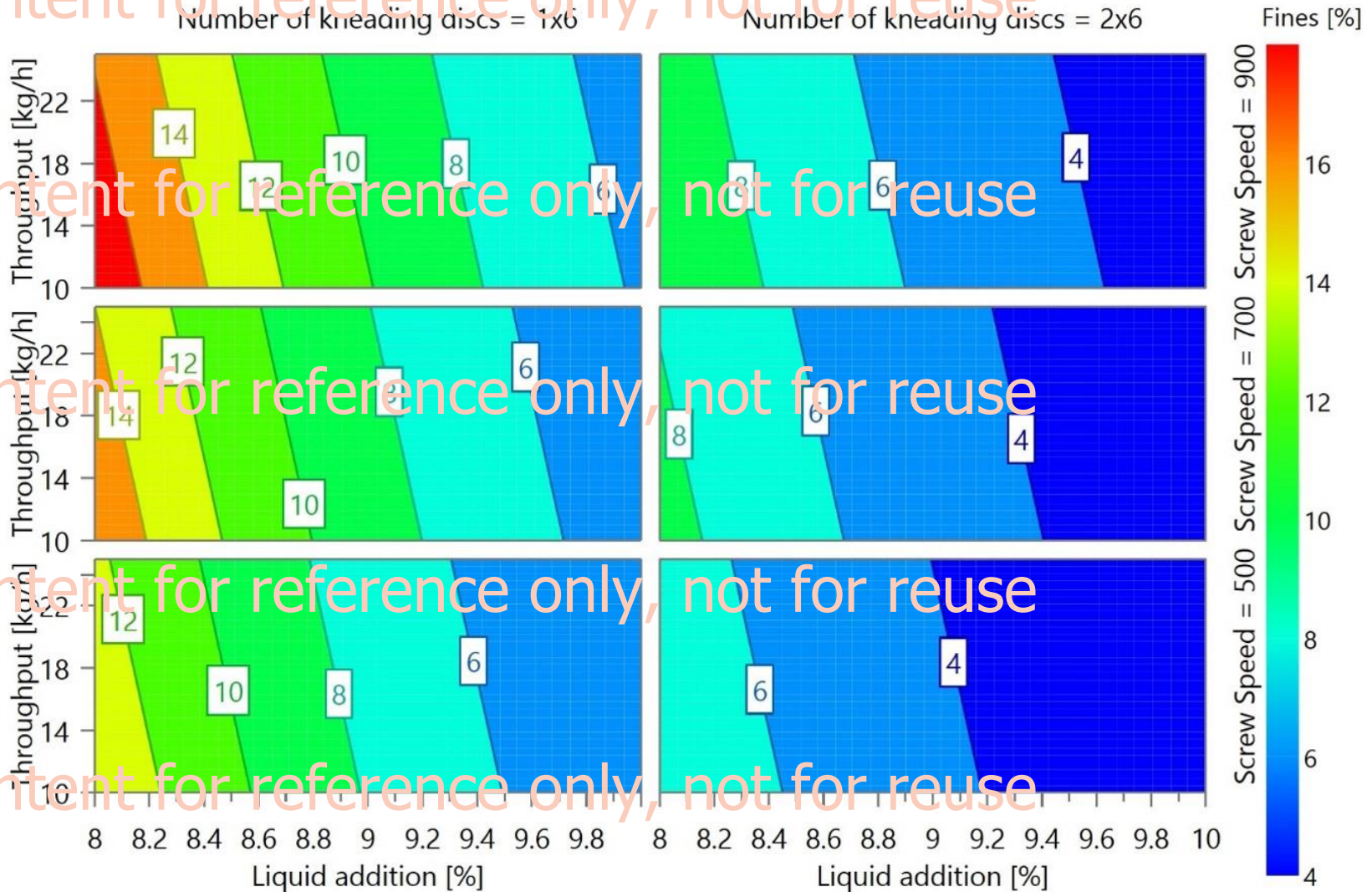
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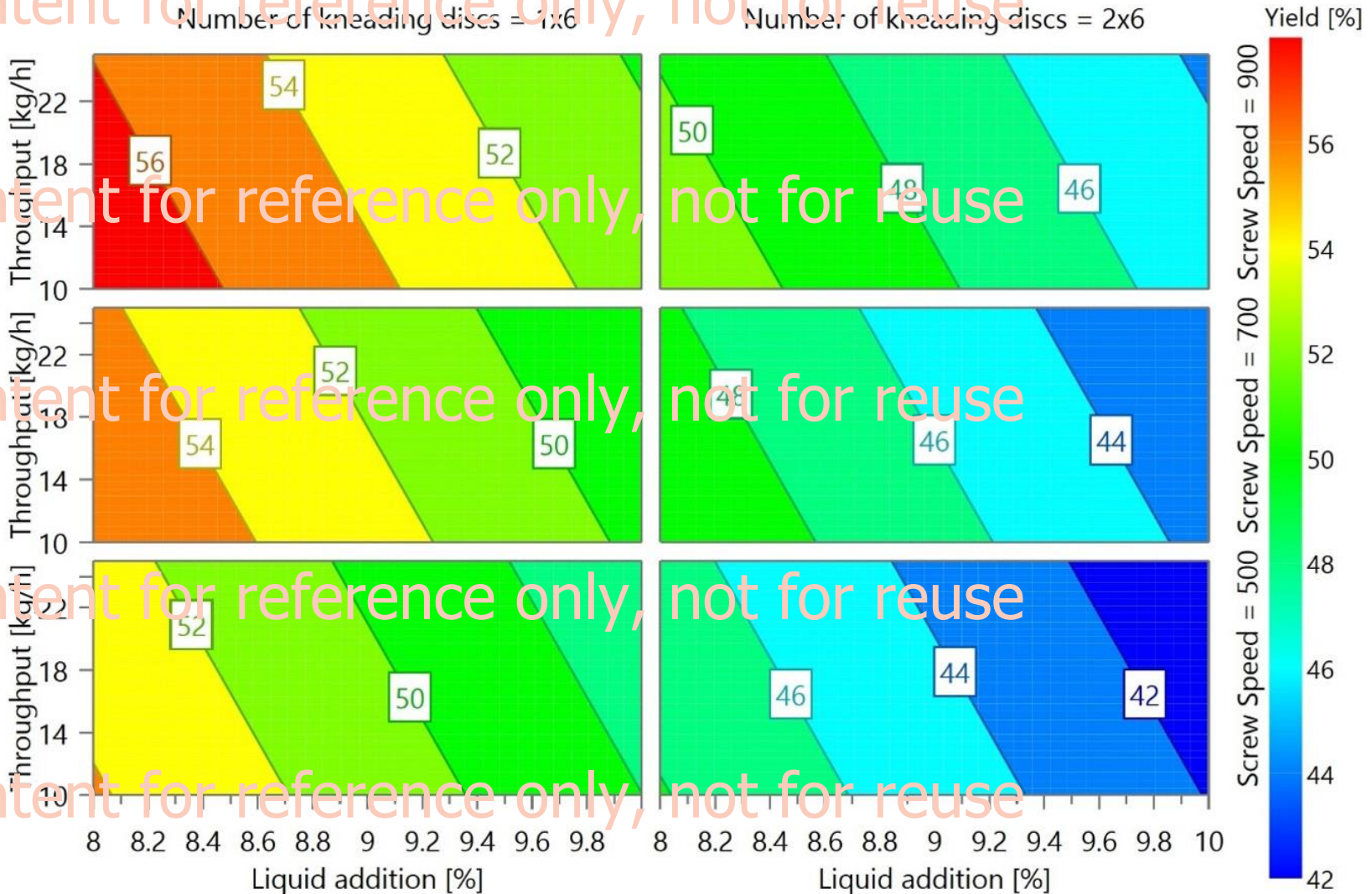
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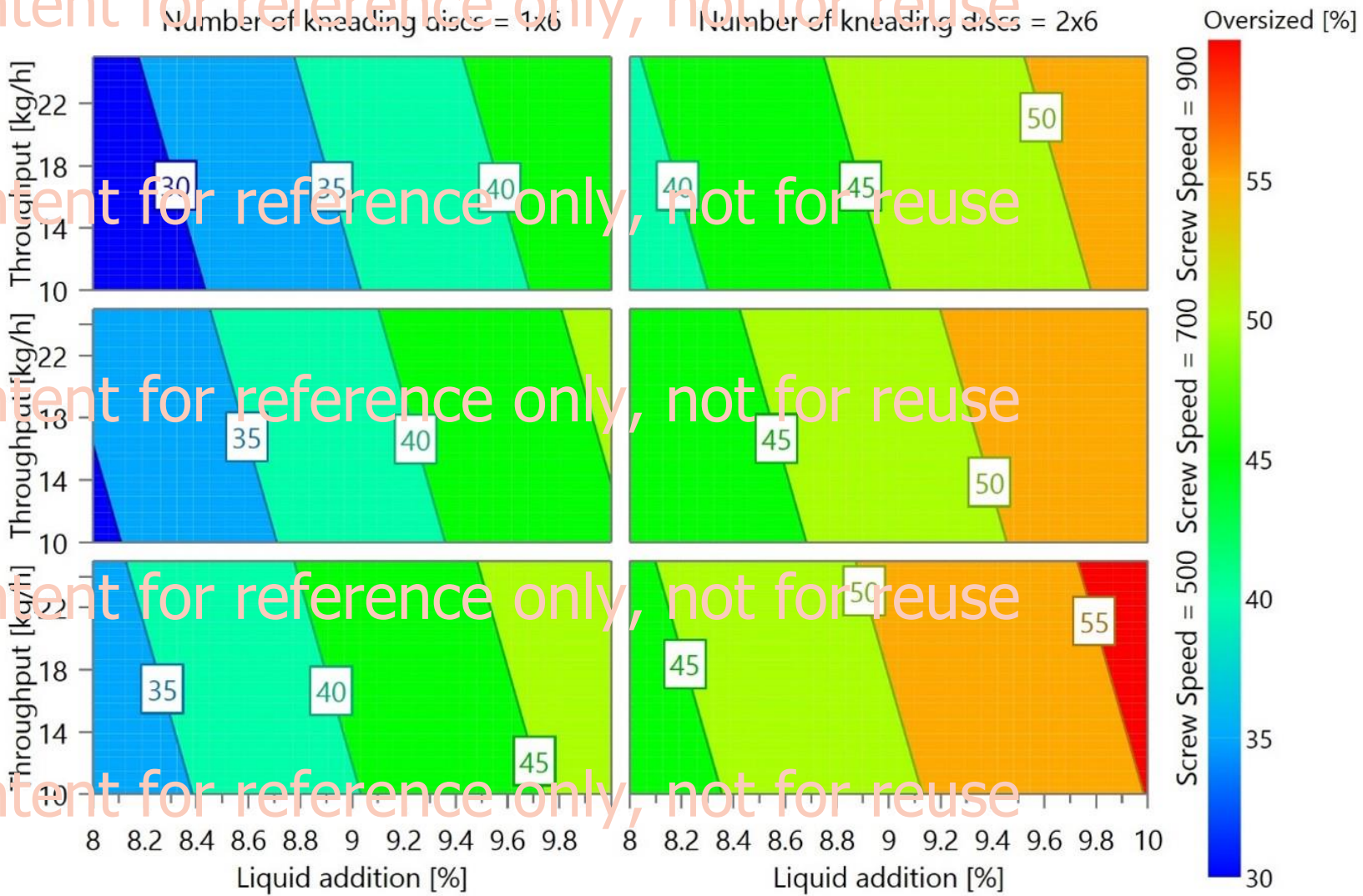
Increasing number of kneading discs and L/S reduced amount of fines



Yield fraction increases at low fill ratio and reduces with increasing L/S

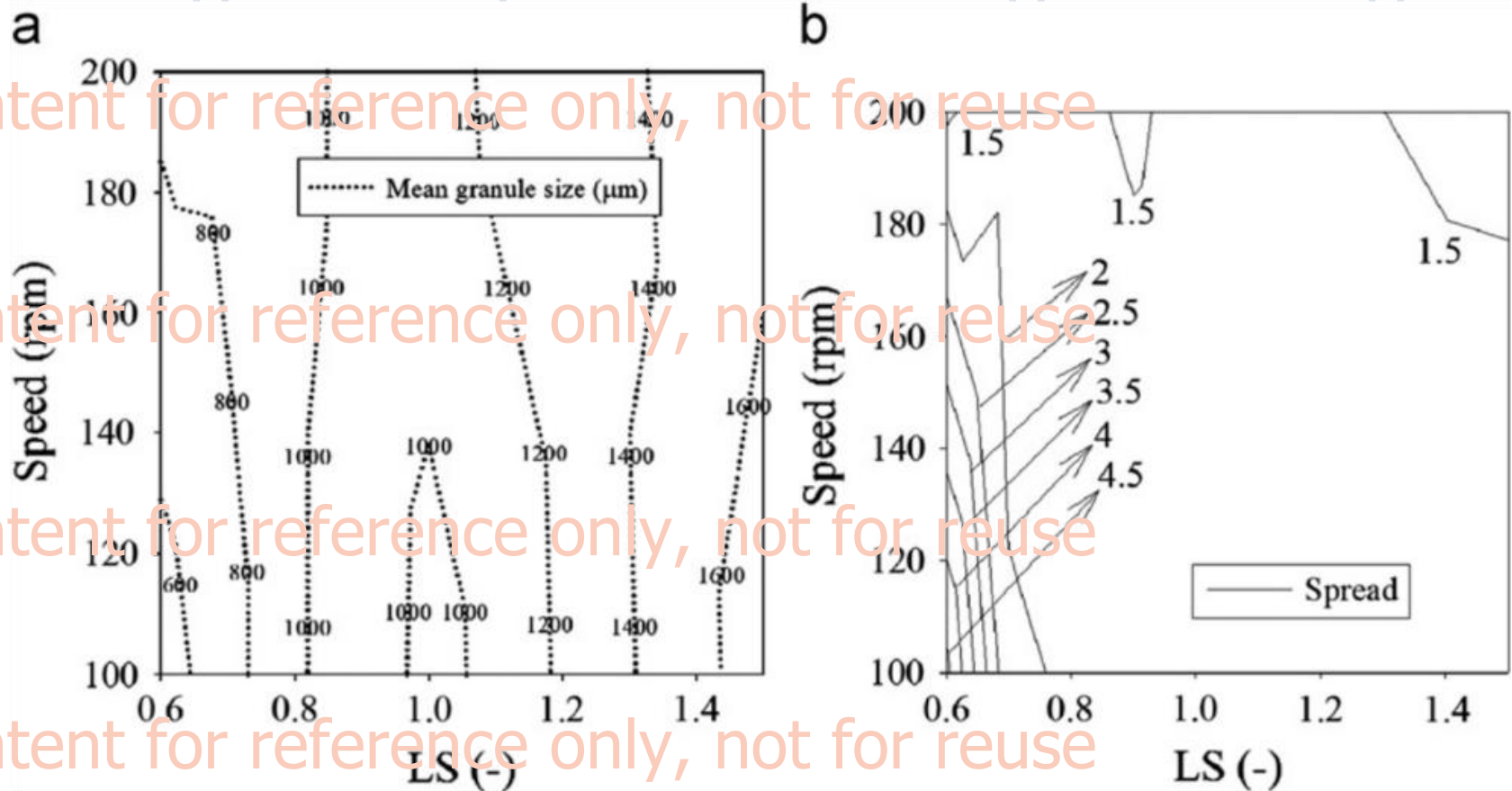


Increasing L/S produced more oversize at high fill ratio



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Regime map: consolidating knowledge



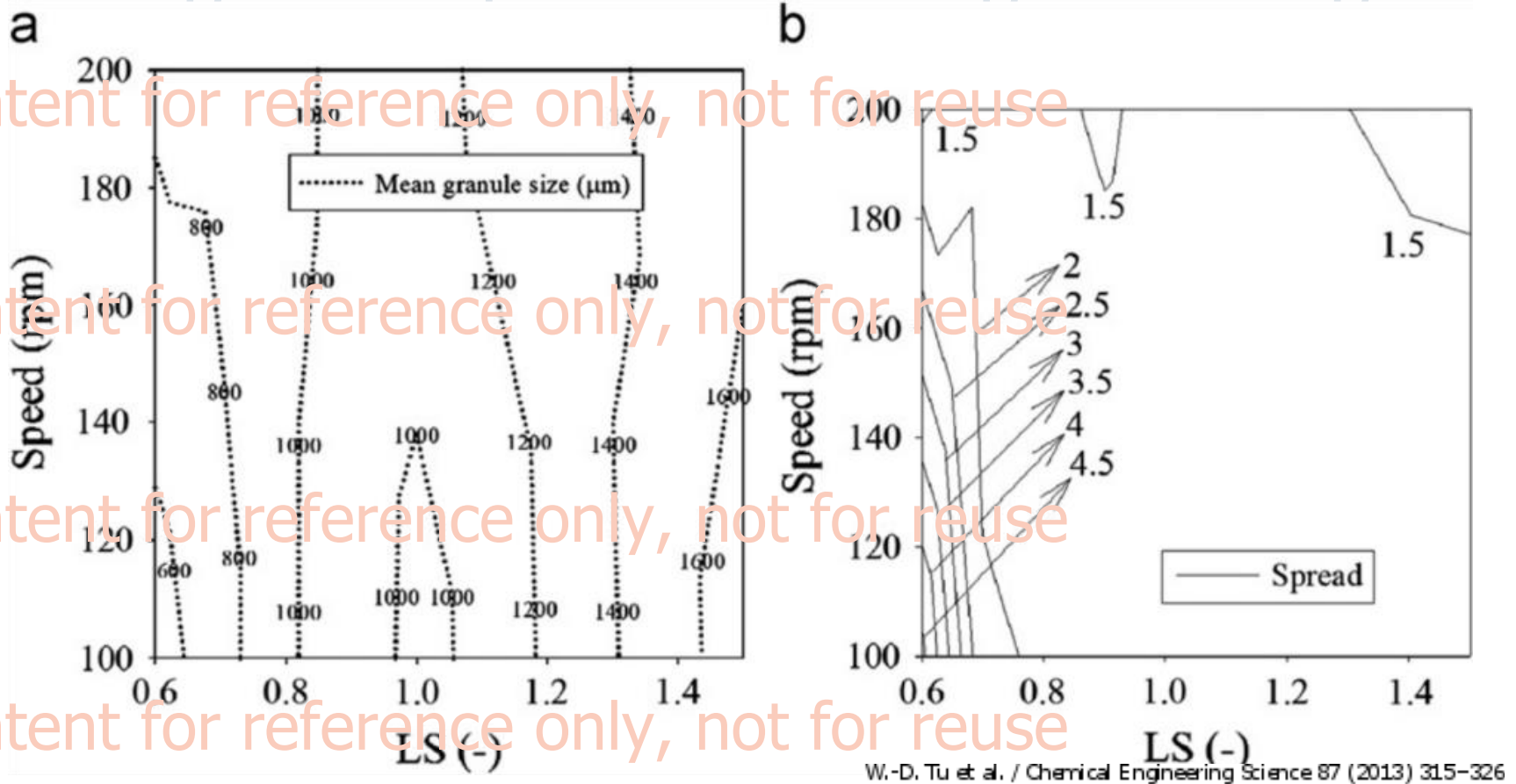
W.-D. Tu et al. / Chemical Engineering Science 87 (2013) 315–326

Boundaries were drawn to separate the growth behavior in terms of the L/S ratio and screw speed.

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Regime map: consolidating knowledge



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Dimensional measurements limits its applicability.

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Growth regime map for wet granulation

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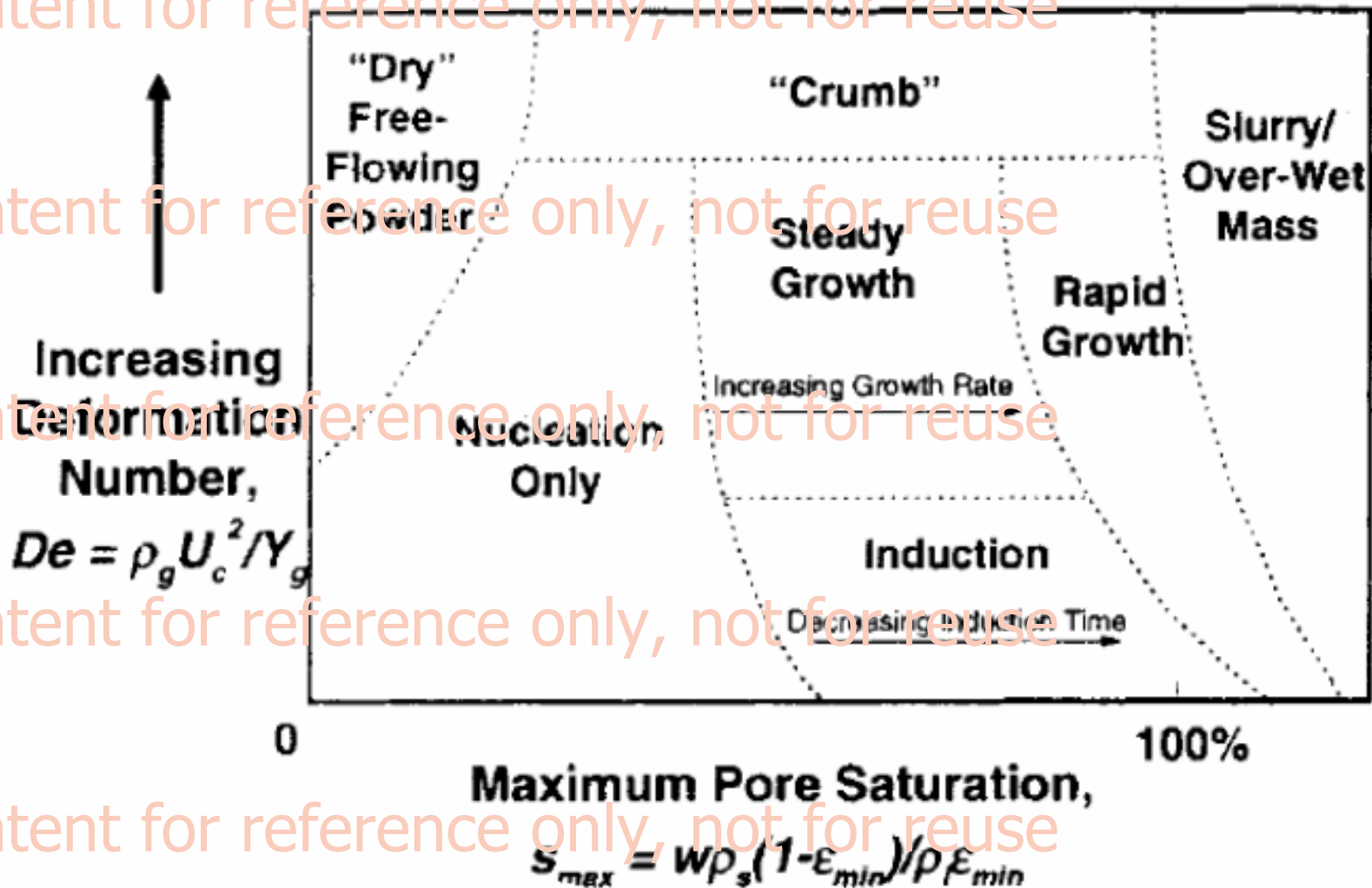
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source: Iveson et al. (1998). AIChE J., 44, 1510 - 1518

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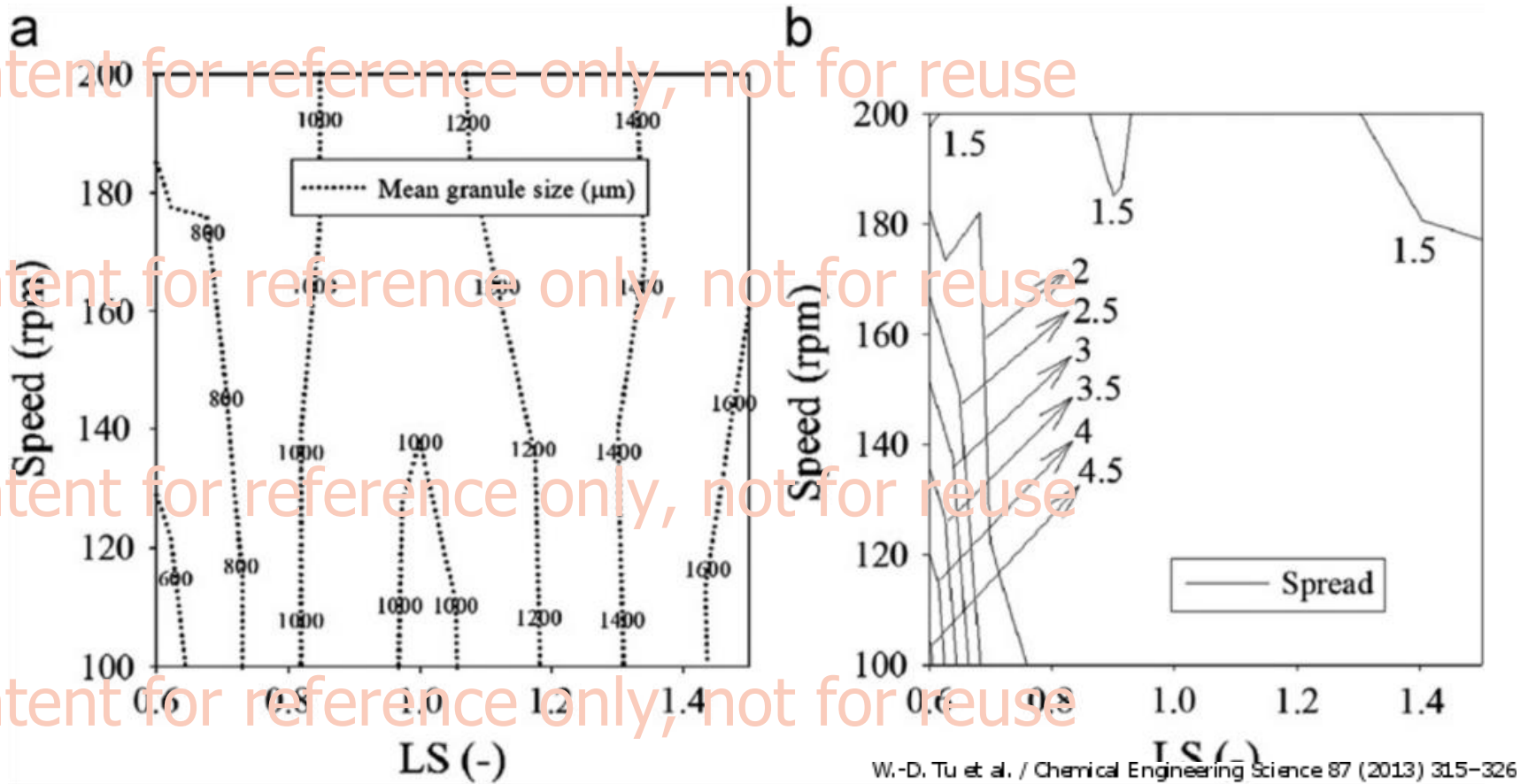
Regime map: consolidating knowledge

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Mean of granule size distribution is not representative.

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Using scale-independent parameter and a broad look on distribution

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Specific mechanical energy (kJ/kg)

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$$SME = \text{motor rating} \times \% \text{ torque} \times \frac{RPM_{oper.}}{RPM_{max.}} \times \frac{\text{gearbox efficiency}(0.97)}{\text{material throughput}}$$

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Liquid to solid ratio (%)

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Quartiles of distribution D25, D50, D75

Quartile ratio D75/ D25

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A higher L/S ratio led to formation of oversized
granules instead of increasing yield

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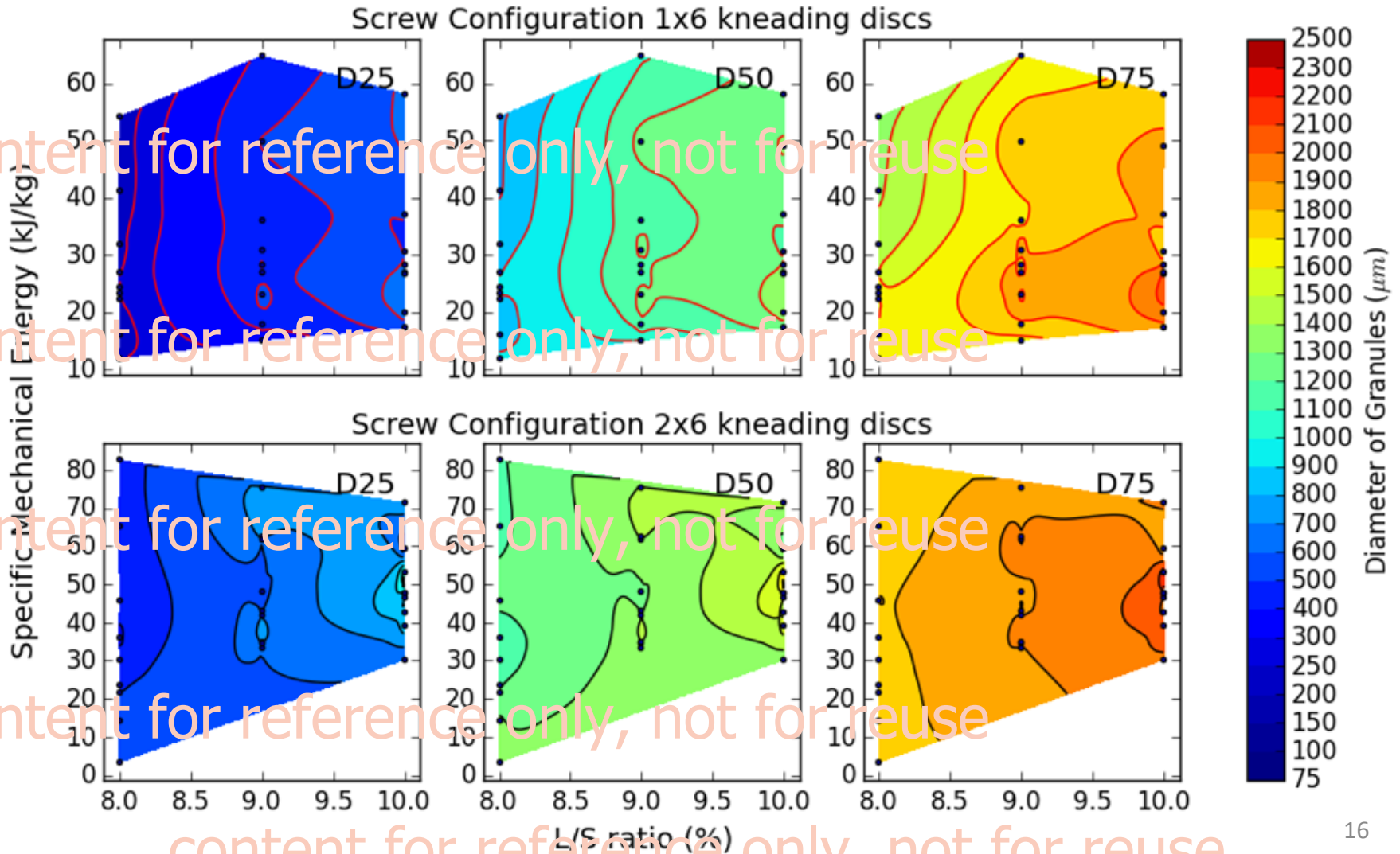
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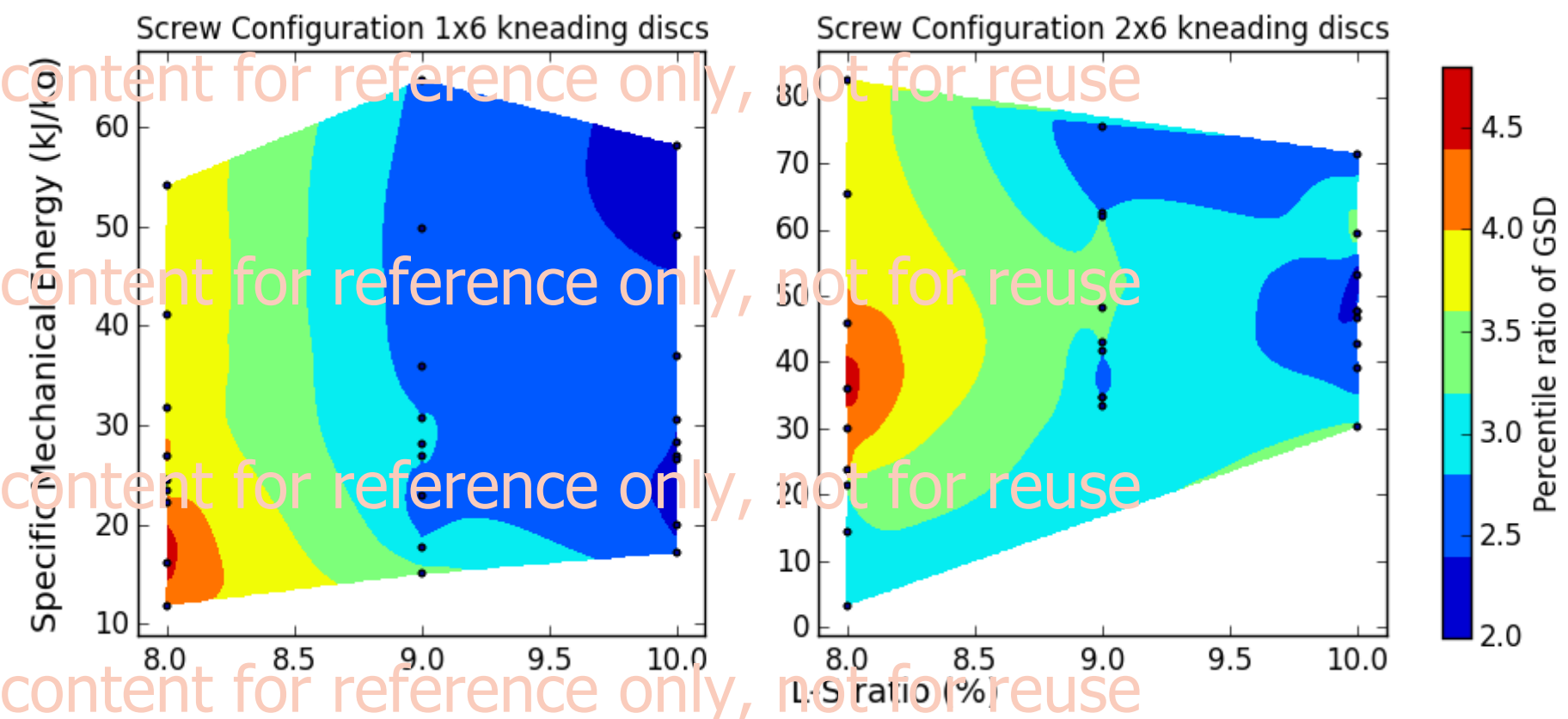
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Width of the distribution reduced at higher L/S ratio and SME level

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Summary

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..the liquid addition should be kept at an intermediate level.

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.. the throughput and screw speed should be increased simultaneously to improve the granulation yield.

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.. more regime maps for other formulations are necessary due to the likely differences in the granulation behavior.

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Perspectives

Formulation properties

$D_p, Y, \gamma \cos \theta \dots$

Process parameter

RPM, P_{input} , shear..

Propose most relevant
dimensionless groups

Generalised
regime map

Aknowledgements

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Laboratory of Pharmaceutical Process Analytical Technology

The logo for BIOMATH. The word 'BIOMATH' is written in a blue, sans-serif font. The letter 'O' is replaced by a circular graphic containing several smaller blue dots of varying sizes, representing a molecular or biological structure.

Model-based analysis and optimization of bioprocesses

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