



The Power of Machine Learning

Image analysis for particle size and shape



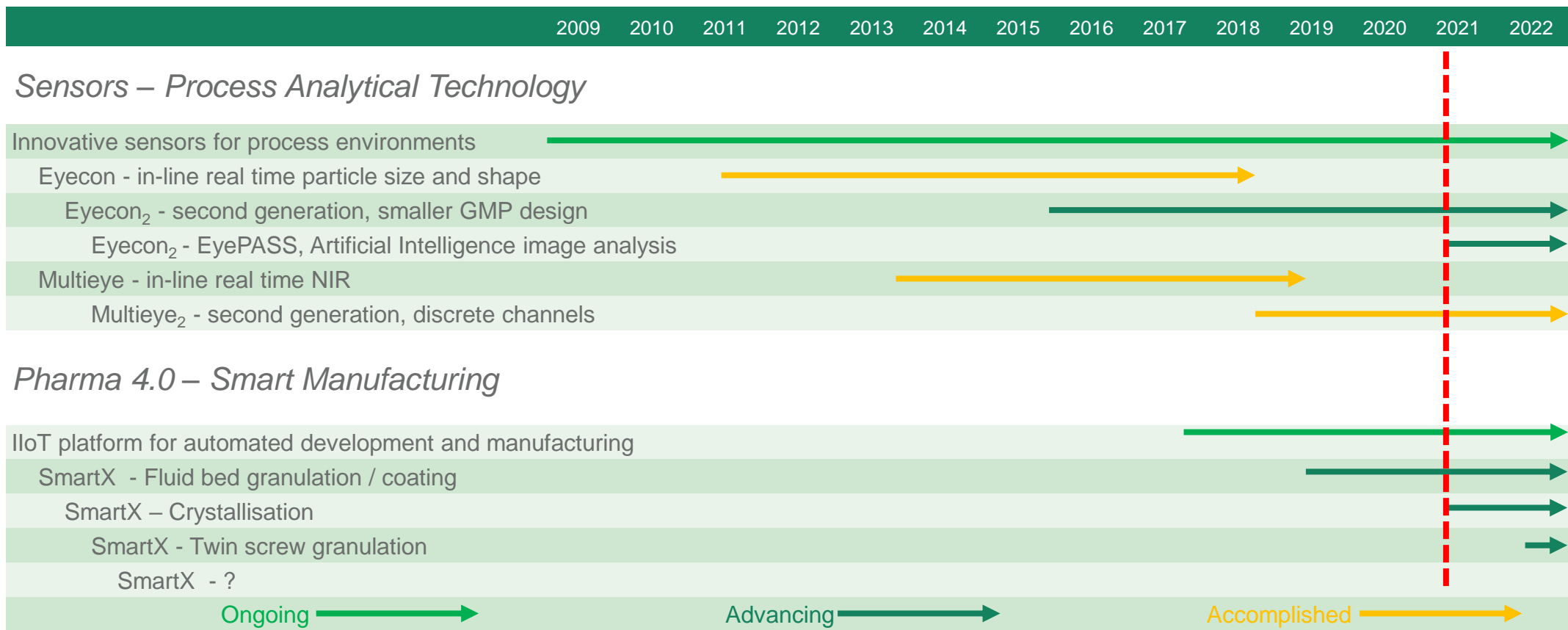
- Innopharma Labs founded 2009
- Expanded into:
 - *Innopharma Education*
 - *Innopharma Technology*
 - *Innopharma Technical Services*
- HQ: Dublin, Ireland
- ~120 employees experienced in STEM, pharma, manufacturing, IT, Software Development



The Journey

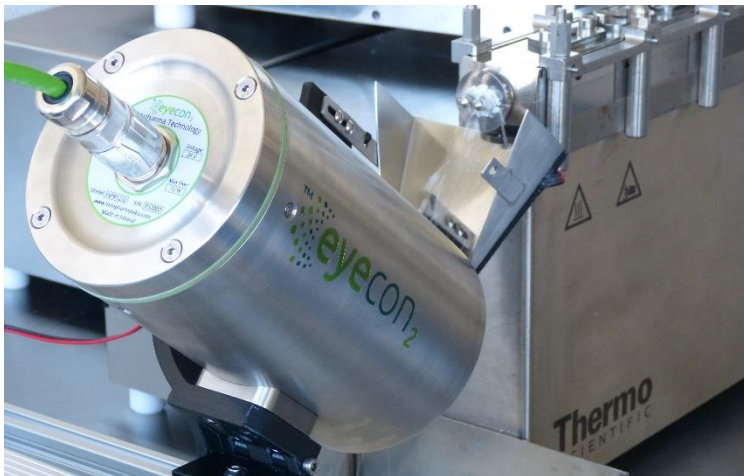


Solutions for Advanced Manufacturing



Eyecon₂ Technical Specifications

Size Range	50 to 5500 µm
Casing materials	304 Stainless Steel, Glass, Silicon (gaskets)
Imaging Area	11.25 x 11.25 mm
Output	PDF. CSV, JPEG full PSD D5-D95.
Instrument Ratings	GMP compliant design EyePASS is both 21 CFR part 11 & GAMP5 Compliant CE Marking ATEX zones 2/22, IP65.
Configurations	In-line and at/offline
Communication	Ethernet and USB OPC UA, OPC DA 3.0

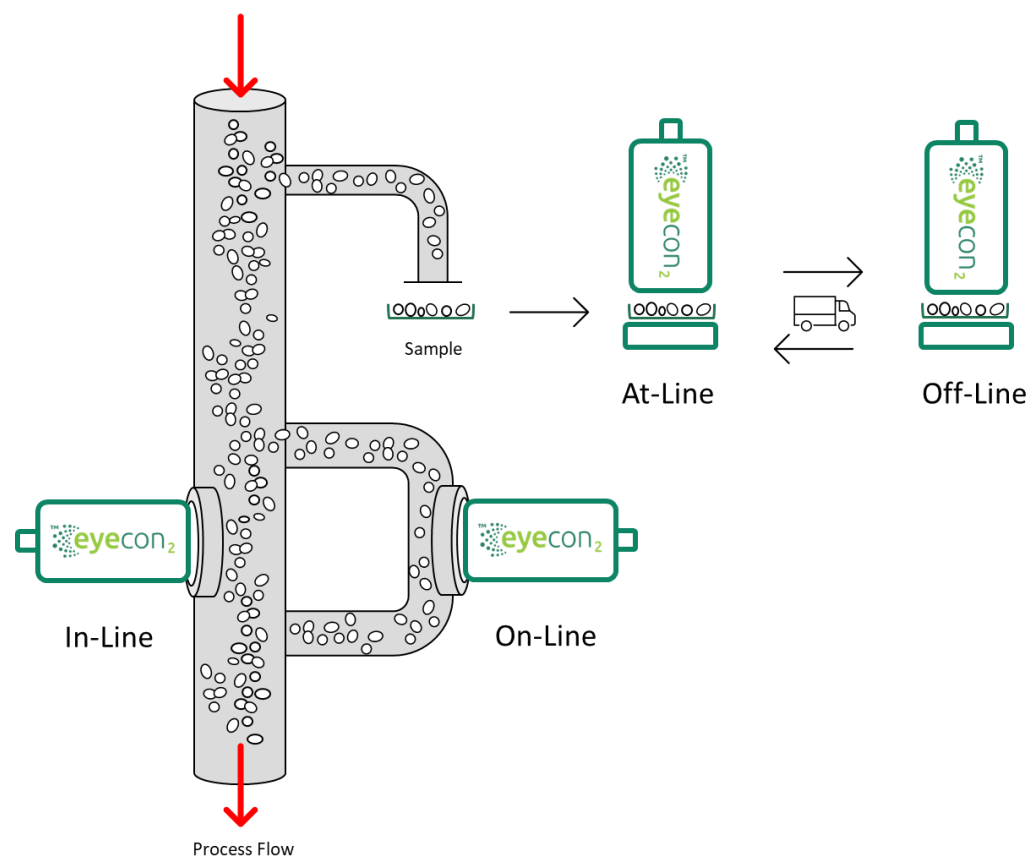


- ✓ *In-line*
- ✓ *At-line*
- ✓ *Non product contact*
- ✓ *Real time*

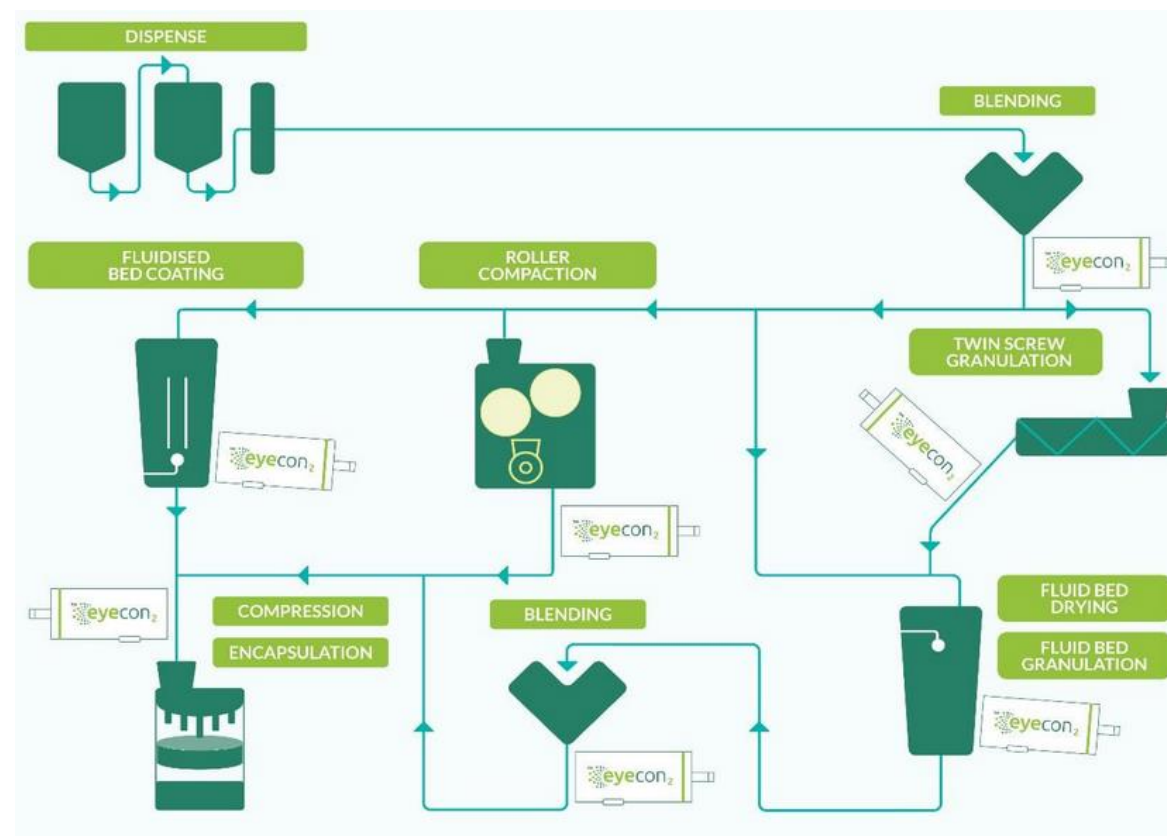


Small particles are Big deal

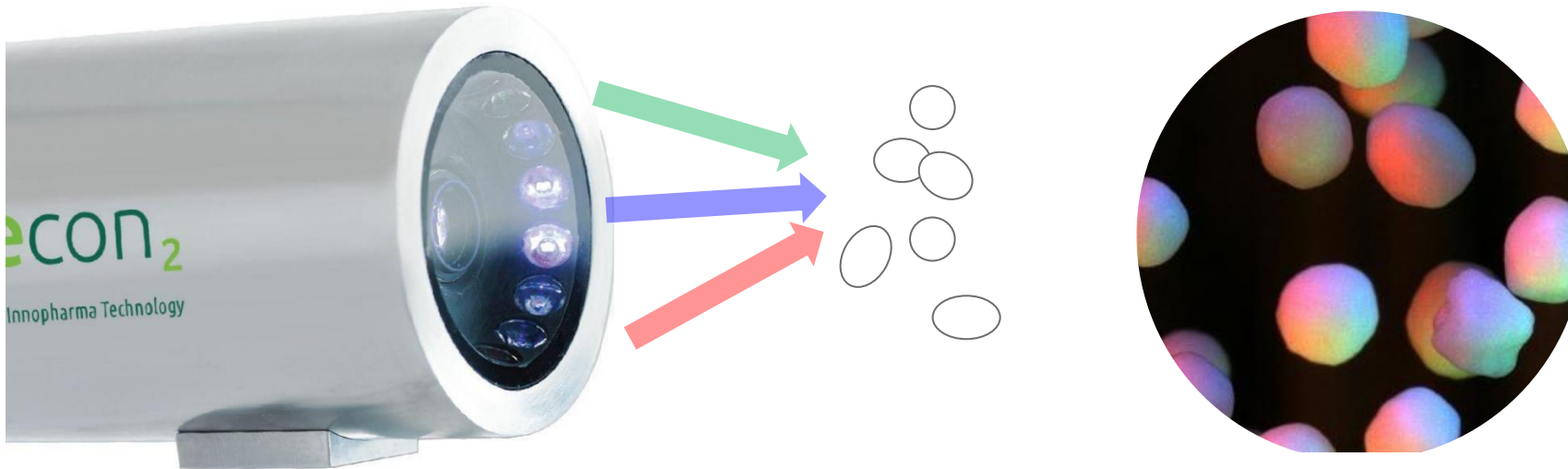
Configurations



Processes



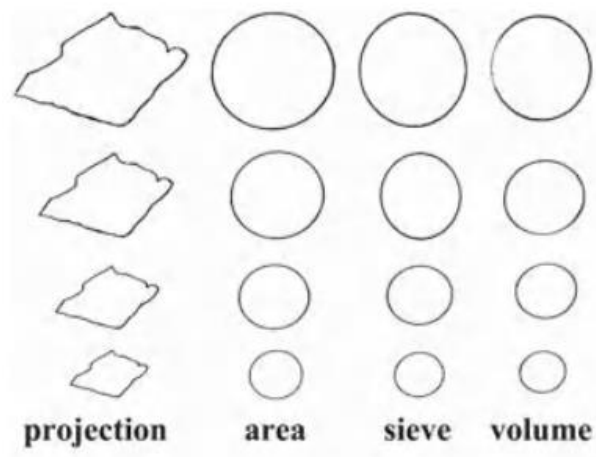
Method of Operation: Image Capture



- A flash-imaging technique is used with an extremely short light-pulse to illuminate moving particles for image capture
- Red, Green and Blue LEDs illuminating the sample from different angles for detection of particle boundaries

Particle size methods

Particle size methods are not created equally, they all have their advantages and disadvantages



2.1 Particle Size

Fig. 2.2 Equivalent sphere concept for arbitrary particles

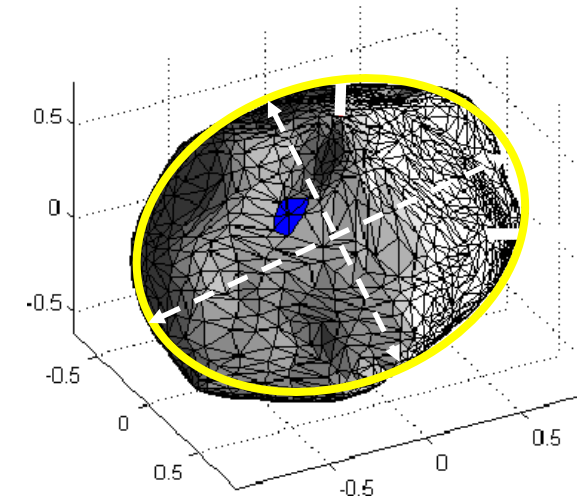
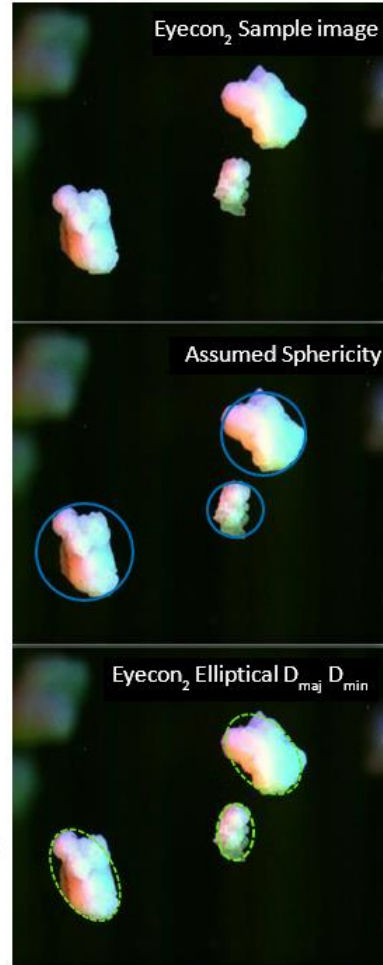
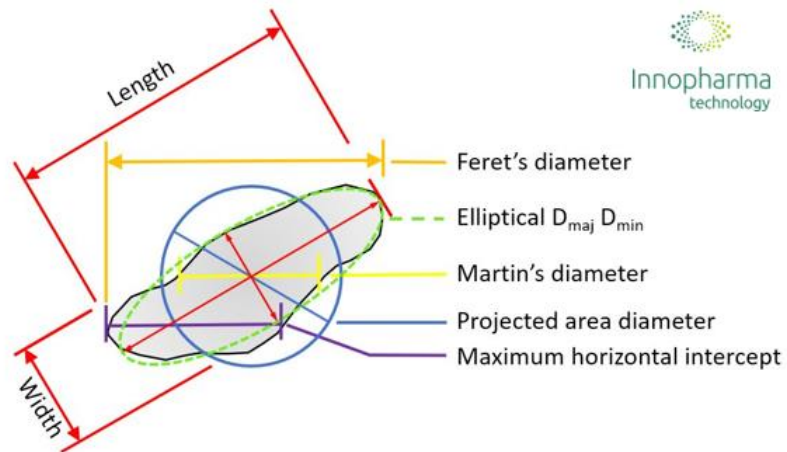
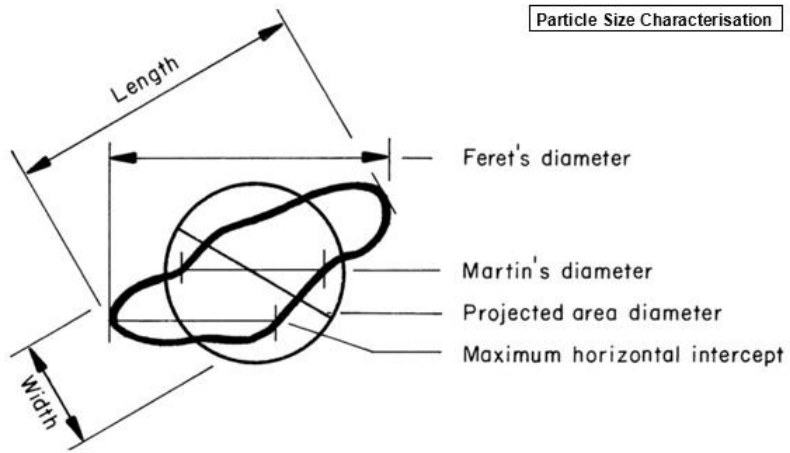
From: Particle Size Measurements, Fundamentals, Practice, Quality, Henk G. Merkus

Horiba Particle Guidebook 2022



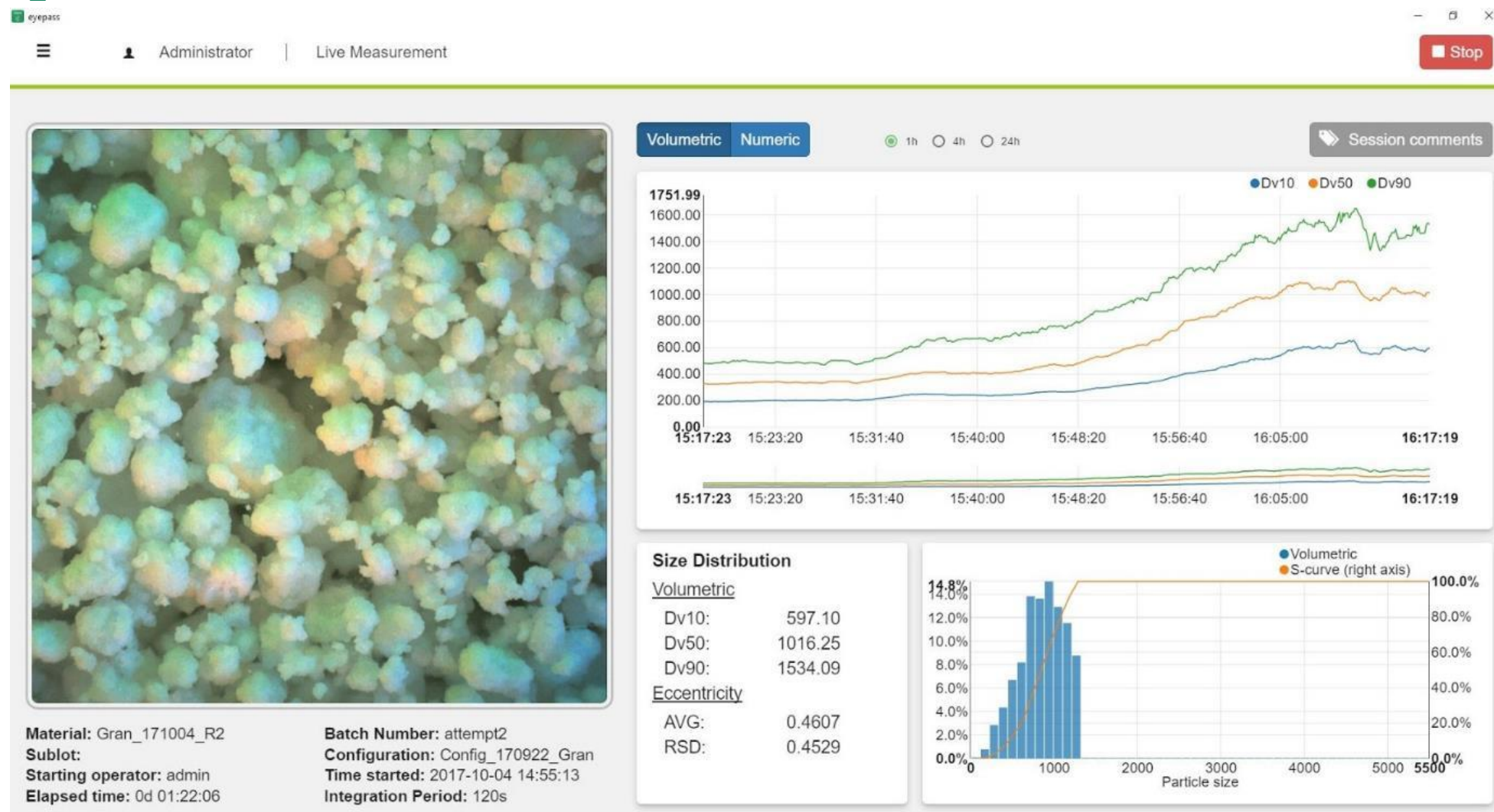
From: https://static.horiba.com/fileadmin/Horiba/Products/Scientific/Particle_Characterization/Particle_Guidebook_2022.pdf

Method of Operation: Image Capture



- Each particle initially identified
- Best-fit ellipse calculated
- Major & minor diameters computed
- PSD/D-values determined

Eyecon₂ Data Output



Eyecon₂ Data Output



Particle Size Report



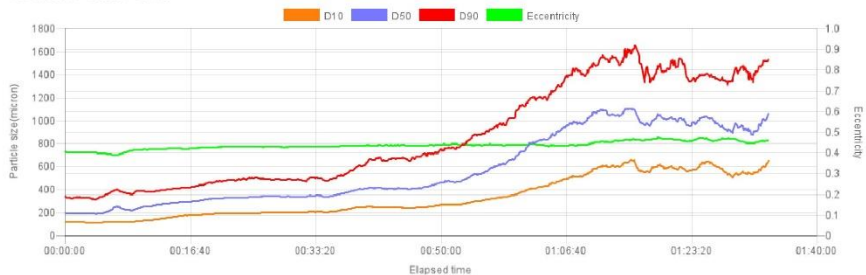
Batch Details

Material	Gran_171004_R2	Configuration	Config_170922_Gran	Start time	2017-10-04 14:55:13
Batch Number	attempt2	Started by	admin	End time	2017-10-04 16:28:59
Sublot		Number of particles	543602	Duration	1:33:47
Instrument details	EYECON2	Eyecon serial number	PS2003	Eyepass version	1.1.4

Alarms & notifications

System events	No
---------------	----

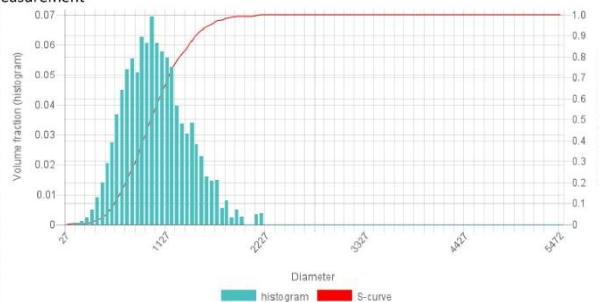
Results: Process Profile



Results: End Point

Data based on final 300 seconds of measurement

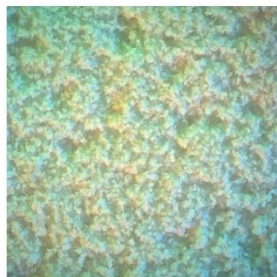
Dv10	582.64
Dv25	752.50
Dv50	981.33
Dv75	1224.60
Dv90	1480.19
Dn10	109.05
Dn25	254.69
Dn50	483.43
Dn75	714.25
Dn90	964.79
Mean	515.82
Median	483.43
Std deviation	326.08
Eccentricity mean	0.4511
Eccentricity RSD	0.4326



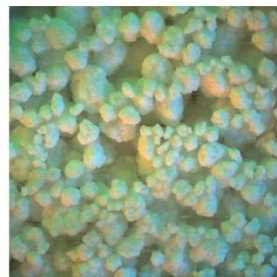
Sample Images



2017-10-04 14:55:14



2017-10-04 15:42:06



2017-10-04 16:28:54

Signature: _____

Date: _____



Innopharma
technology

Eyecon₂ Data Output



Eyecon Particle Size Report - Innopharma Technology																																																		
Batch Details																																																		
Material	Batch Number	Configuration	Started by	Start date	Start time	End date	End time	Duration (seconds)																																										
Gran_171004	attempt_1	n/a	Config_1 admin	2017-10-	14:55:13	2017-10-	16:28:59	5626.06																																										
Instrument Details																																																		
Model	Serial number	EyePASS version																																																
EYECON2	PS2003	1.1.4																																																
Configuration Details																																																		
Name	Mode	Image size	Short duration	Integration	Min. Ref	Max. Ref	Max. det	False avg	Edge correction	Analysis	Material	Alarms enabled	Report type	Product type	All camera	R camera	G camera	B camera	R LED gain	G LED gain	B LED gain	Focus position (Eyecon 2 only)																												
Config_17092	In Line	12	0	120	25	5000	1400	0.25	60	301	0	false	Default	300	53	100	40	75	60	50	251																													
End Point Results - calculated from final 300 seconds of process																																																		
			D_v10	D_v50	D_v90	D_n10	D_n50	D_n90	Mean diameter	Median	Std deviation	d1 average	d2 average	Shape factor	Shape factor	5% vol	10% vol	15% vol	20% vol	25% vol	30% vol	35% vol	40% vol	45% vol	50% vol	55% vol	60% vol	65% vol	70% vol	75% vol	80% vol	85% vol	90% vol	95% vol	100% vol															
			582.64	981.33	1480.19	109.05	483.43	964.79	515.82	483.43	326.08	553.18	478.46	0.4511	0.4326	491.25	582.64	649.26	700.41	752.5	802.81	854.17	896.13	938.96	981.33	1022.46	1069.56	1121.73	1169.32	1224.6	1303.89	1400.34	1480.19	1616.77	2183.39															
Process Profile - all results based on integration period of 120 seconds																																																		
Date	Time	Elapsed	D_v10	D_v50	D_v90	D_n10	D_n50	D_n90	Mean diameter	Median	Std deviation	d1 average	d2 average	Shape factor	Shape factor	5% vol	10% vol	15% vol	20% vol	25% vol	30% vol	35% vol	40% vol	45% vol	50% vol	55% vol	60% vol	65% vol	70% vol	75% vol	80% vol	85% vol	90% vol	95% vol	100% vol															
04/10/2017	14:55:14	0.63	115.43	197.14	340.44	107.46	124.48	203.48	141.31	124.48	46.69	149.11	133.51	0.4027	0.3867	110.98	115.43	120.62	126.04	132.02	139.79	150.15	178.79	189.88	197.14	204.67	216.19	227.27	246.14	269.78	290.04	314.68	340.44	366.5	469.02															
04/10/2017	14:55:22	8.45	115.45	195.99	331.33	107.62	124.53	203.29	141.15	124.53	45.74	148.98	133.31	0.4039	0.3853	111	115.45	120.55	125.89	131.59	138.04	147.85	175.21	188.15	195.99	203.27	212.92	225.5	238.21	261.21	283.12	303.03	331.33	361.31	469.02															
04/10/2017	14:55:29	16.23	115.44	195.96	323.75	107.72	124.37	203.84	140.96	124.37	45.34	148.65	133.27	0.4013	0.3828	110.83	115.44	120.37	125.51	130.96	137.76	147.27	169.1	187.45	195.96	203.6	212.2	222.89	236.55	260.29	280.66	297.85	323.75	358.87	469.02															
04/10/2017	14:55:37	24.02	115.75	196.7	325.89	107.93	124.55	204.71	141.3	124.55	45.54	149.01	133.58	0.4008	0.384	111.04	115.75	120.61	125.82	131.45	138.13	147.83	171.39	188.45	196.7	203.98	213.01	223.14	237.7	261.2	279.73	298.36	325.89	361.73	469.02															
04/10/2017	14:55:45	31.83	115.67	196.65	324.43	107.99	124.46	204.75	141.25	124.46	45.5	148.97	133.54	0.4011	0.384	111.08	115.67	120.44	125.69	131.32	138.04	147.87	171.4	188.38	196.65	203.98	212.69	223.28	236.88	260.85	279.22	297.57	324.43	364.11	469.02															
04/10/2017	14:55:53	39.64	115.69	196.38	326.19	108.1	124.43	204.67	141.17	124.43	45.43	148.88	133.45	0.4011	0.3842	111.11	115.69	120.45	125.59	131.04	137.84	147.56	170.07	187.77	196.38	203.94	212.55	223.11	236.73	260.98	279.94	298.61	326.19	365.81	469.02															
04/10/2017	14:56:01	47.44	115.57	195.5	323.55	108.08	124.54	204.29	140.99	124.54	45.07	148.7	133.28	0.4012	0.3847	111.01	115.57	120.4	125.51	130.87	137.55	146.64	166.85	187.09	195.5	203.37	211.46	221.64	234.66	258.38	278.44	296.3	323.55	364	469.02															
04/10/2017	14:56:08	55.24	115.57	195.35	323.5	108.05	124.54	204.13	140.97	124.54	45.06	148.67	133.27	0.4011	0.3846	110.99	115.57	120.38	125.5	130.86	137.49	146.49	166.73	186.86	195.35	203.19	211.33	221.6	235.24	258.91	278.82	296.22	323.5	364.66	469.02															
04/10/2017	14:56:16	63.04	115.6	195.18	322.5	108.01	124.57	203.94	140.94	124.57	45.01	148.67	133.22	0.4018	0.3842	110.98	115.6	120.41	125.47	130.85	137.35	146.33	166.17	186.77	195.18	203.08	211.16	221.43	234.71	259.11	278.97	296.33	322.5	365.17	469.02															
04/10/2017	14:56:24	70.85	115.64	195.12	324.22	107.99	124.58	203.75	140.88	124.58	45.03	148.59	133.17	0.4016	0.3845	110.98	115.64	120.42	125.51	130.83	137.27	146.35	166.22	186.7	195.12	203.05	211.23	221.46	234.53	259.46	279.02	296.91	324.22	367.26	541.35															
04/10/2017	14:56:32	78.66	115.63	195.34	326.43	107.92	124.53	203.75	140.85	124.53	45.1	148.56	133.13	0.4015	0.3853	110.95	115.63	120.42	125.46	130.84	137.27	146.49	166.38	186.72	195.34	203.15	211.39	221.56	234.69	260.13	279.69	298.13	326.43	370.49	541.35															
04/10/2017	14:56:40	86.47	115.61	195.35	329.01	107.93	124.5	203.62	140.8	124.5	45.1	148.52	133.09	0.4017	0.3856	110.98	115.61	120.41	125.43	130.8	137.25	146.4	166	186.65	195.35	203.03	211.29	221.51	234.87	260.29	280.4	298.86	329.01	371.2	541.35															
04/10/2017	14:56:47	94.25	115.56	195.32	331.33	107.91	124.45	203.48	140.72	124.45	45.07	148.43	133.01	0.4015	0.3859	110.94	115.56	120.36	125.35	130.74	137.15	146.31	165.46	186.48	195.32	203.02	211.26	221.52	235	260.29	280.58	298.86	331.33	371.98	541.35															
04/10/2017	14:56:55	102.06	115.52	194.99	329.47	107.88	124.42	203.41	140.65	124.42	44.95	148.37	132.94	0.4017	0.3864	110.93	115.52	120.28	125.26	130.67	137	146.06	164.55	186.33	194.99	202.79	210.8	220.99	234.32	259.63	279.82	297.5	329.47	370.68	541.35															
04/10/2017	14:57:03	109.88	115.55	195.2	330.58	107.89	124.43	203.59	140.7	124.43	45.02	148.42	132.99	0.4017	0.3864	110.97	115.55	120.31	125.31	130.74	137.15	146.24	164.73	186.42	195.2	203.01	211.02	221.3	234.58	259.84	280.28	298.76	330.58	371.7	541.35															
04/10/2017	14:57:11	117.68	115.54	195.11	331.45	107.88	124.41	203.67	140.67	124.41	44.95	148.36	132.95	0.4016	0.3865	110.97	115.54	120.28	125.27	130.7	137.08	146.03	164.28	186.33	195.11	203.02	211	221.02	234	259.14	279.81	298.41	331.45	372.01	541.35															
04/10/2017	14:57:19	125.47	115.53	194.82	329.48	107.92	124.36	203.66	140.55	124.36	44.75	148.24	132.87	0.4014	0.3862	110.96	115.53	120.22	125.12	130.5	136.75	145.46	162.59	185.78	194.82	202.77	210.63	220.32	232.72	257.2	278.99	296.4	329.48	371.6	541.35															
04/10/2017	14:57:26	133.27	115.49	194.6	329.79	107.96	124.31	203.41	140.41	124.31	44.6	148.07	132.74	0.4012	0.3865	110.94	115.49	120.12	124.96	130.32	136.48	144.9	161.17	185.2	194.6	202.39	210.31	219.81	232.1	255.92	278.36	296.39	329.79	372.31	541.35															
04/10/2017	14:57:34	141.07	115.44	194.2	329.59	107.97	124.3	203.02	140.25	124.3	44.39	147.92	132.59	0.4013	0.3869	110.96	115.44	120.02	124.86	130.12	136.14	144.45	159.58	184.66	194.2	201.91	209.83	219.08	231.24	254.38	277.3	296.03	329.59	372.86	541.35															
04/10/2017	14:57:42	148.88	115.3	193.56	326.7	107.88	124.16	202.6	140.02	124.16	44.21	147.68	132.37	0.4014	0.3869	110.89	115.3	119.89	124.64	129.86	135.79	143.95	158.21	184	193.56	201.57	209.45	218.71	230.74	253.47	277.11	295.8	326.7	372.27	541.35															
04/10/2017	14:57:50	156.68	115.29	193.19	326.44	107.9	124.16	202.52	139.9	124.16	44.07	147.55	132.2																																					



EyePASS Version 3.0

Machine learning image analysis

EyePASS

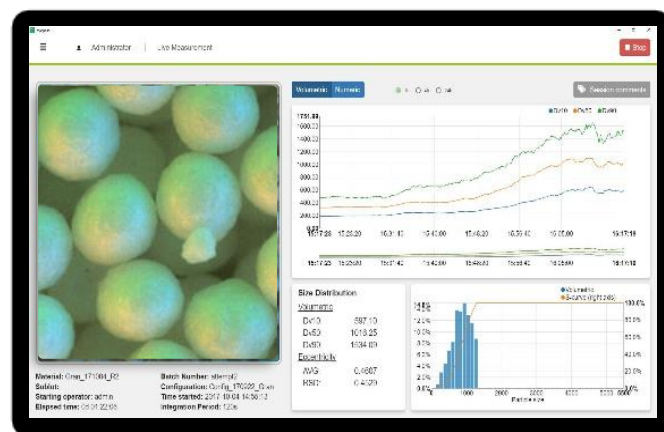


Image Analysis

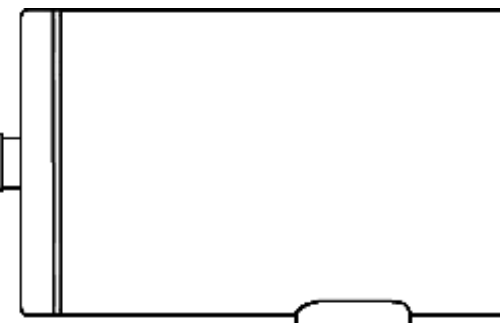
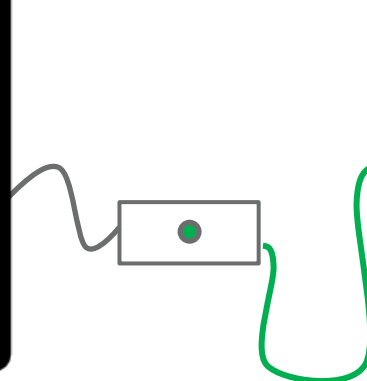
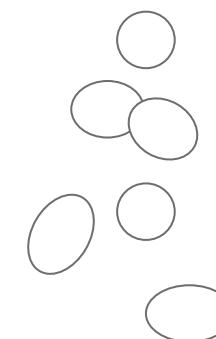
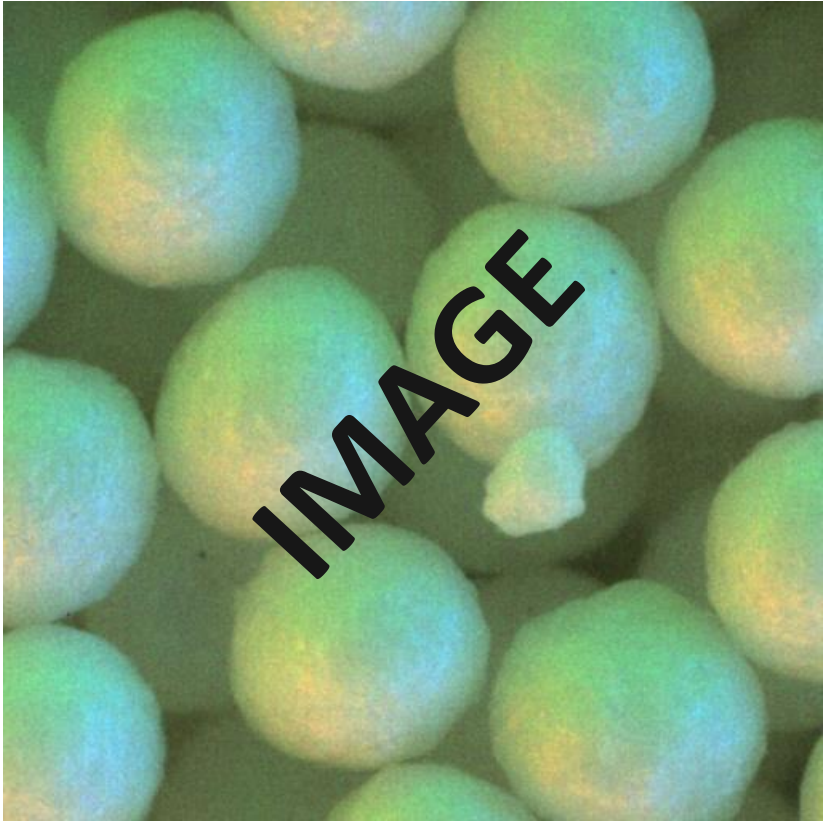


Image Acquisition



• • • • •

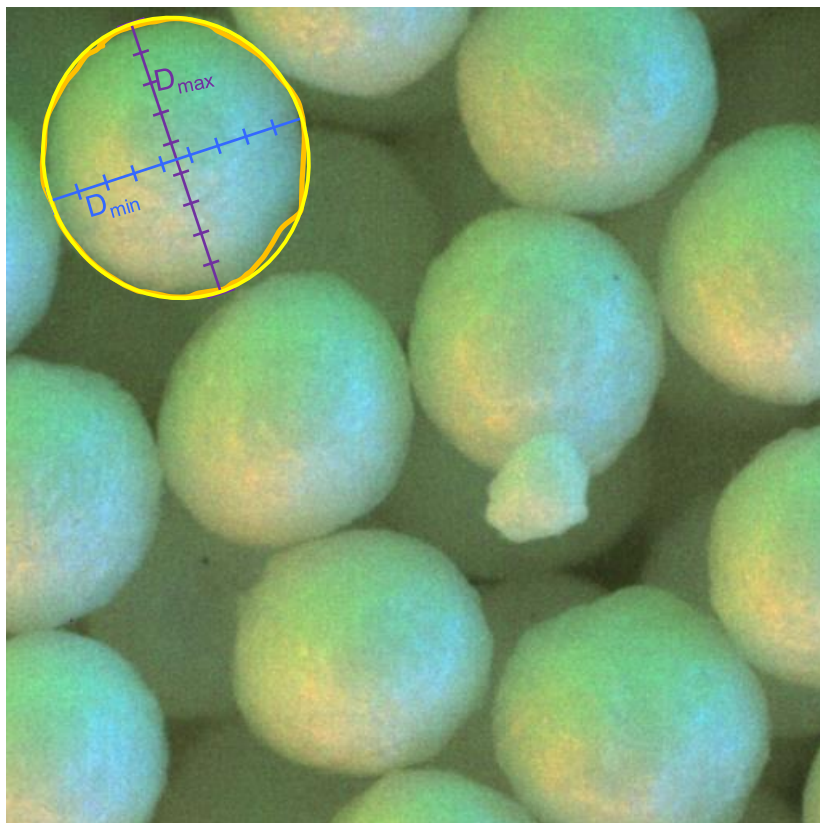
What is Image Analysis?



Analysis

D _{min}	D _{max}
1046.2	1078.89
1061.42	1138.09
1147.19	1336.78
1659.39	1987.52
840.353	1721.95
1065.67	1133.62
983.273	1140.41
1049.12	1151.69
943.567	1142.3
980.434	1178.46
914.148	1095.72
885.818	1085.85

What does it involve?



Vision Task

Delineate objects of interest
e.g., particles



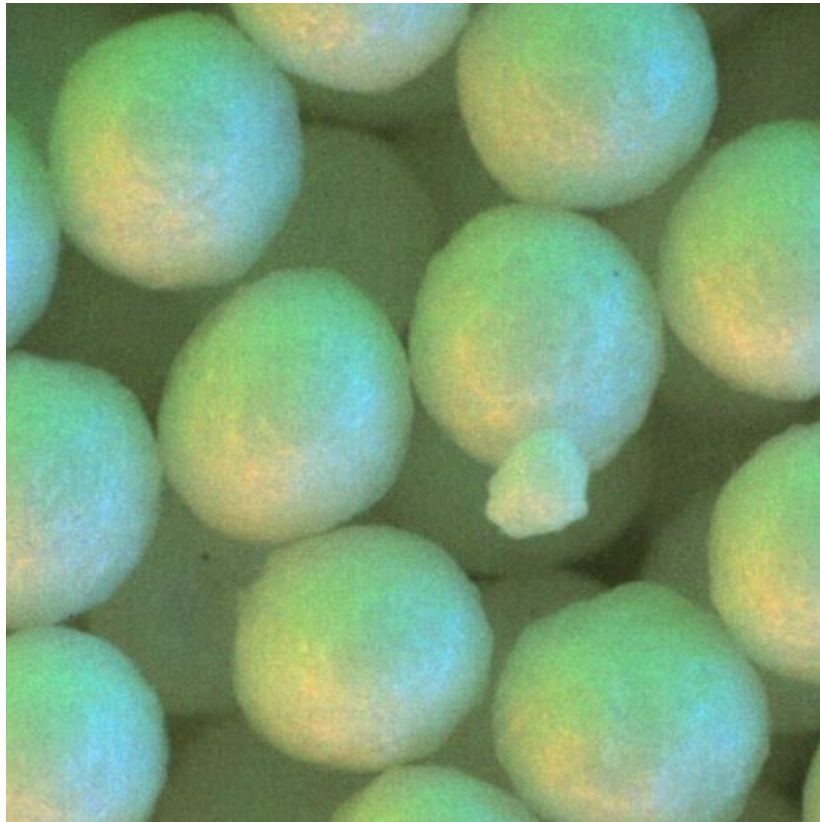
Measurement Task

Measure necessary attributes of each object
e.g., major/minor axes lengths



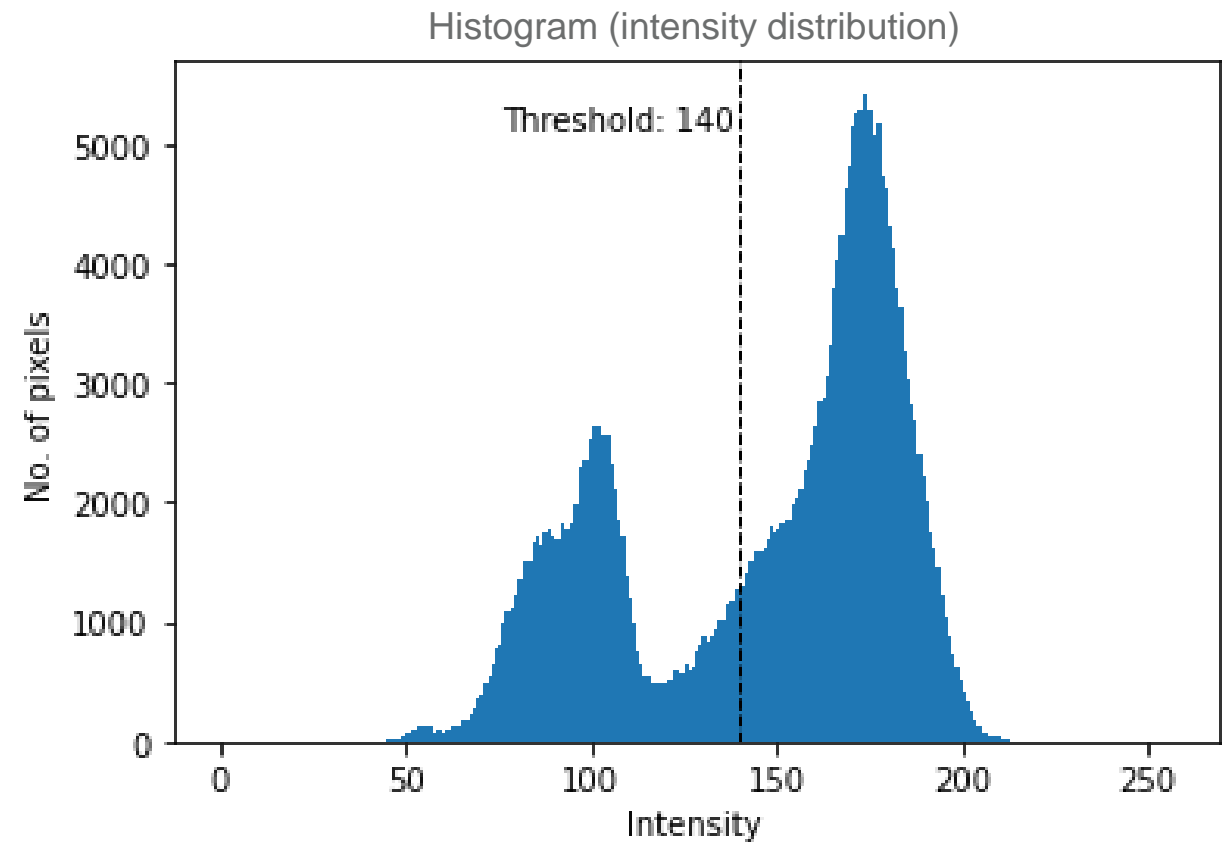
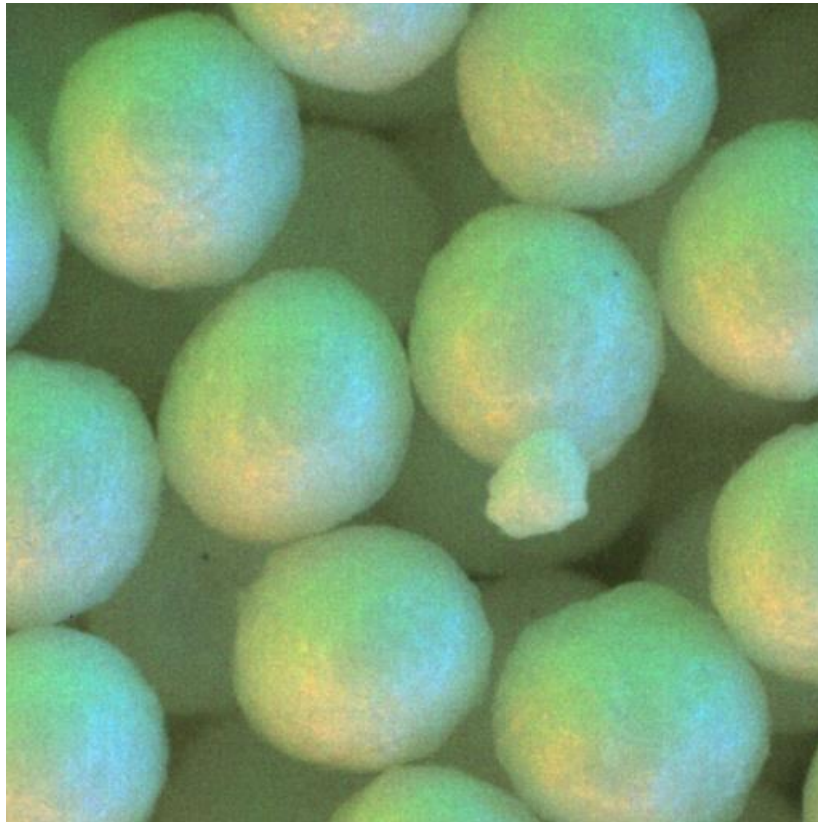
• • • • •

Image Segmentation



The goal as defined by a human observer

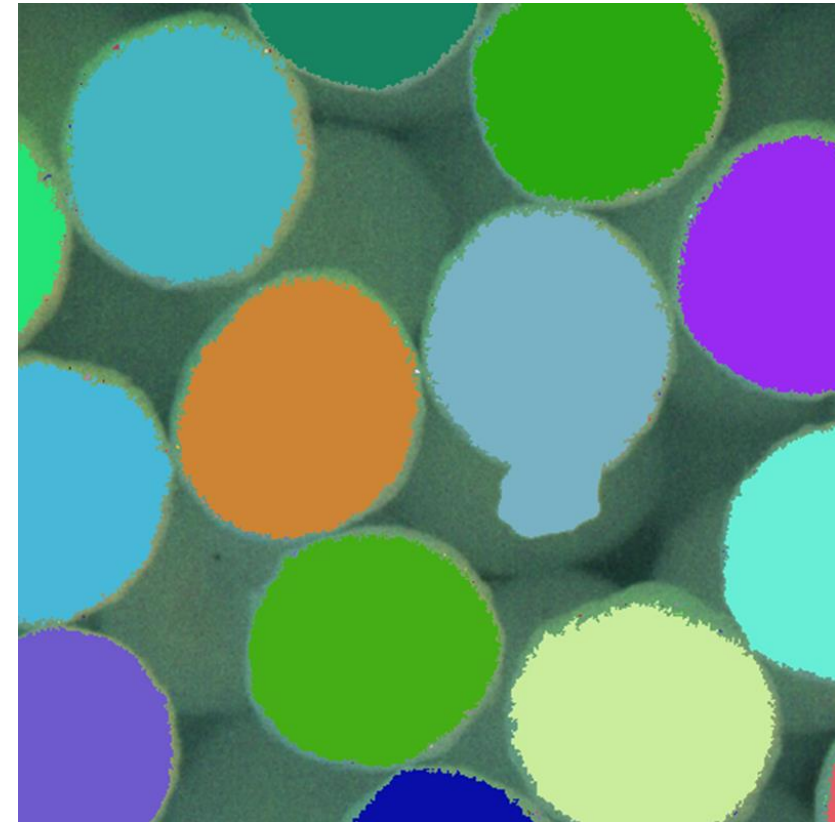
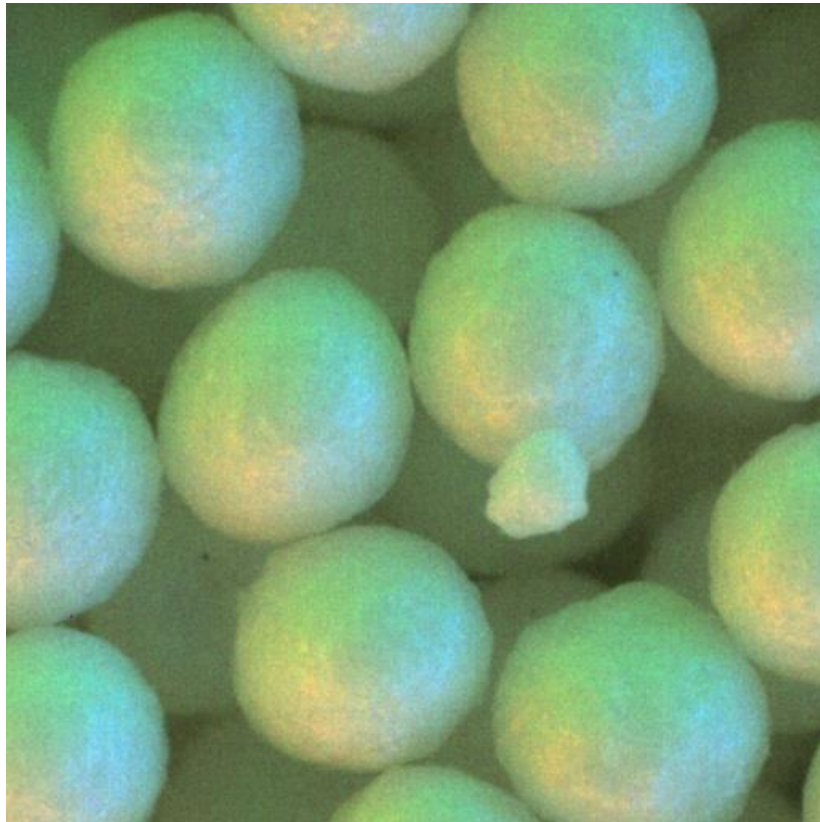
Image Segmentation



Classical Image Processing - Thresholding

-
-
-
-
-

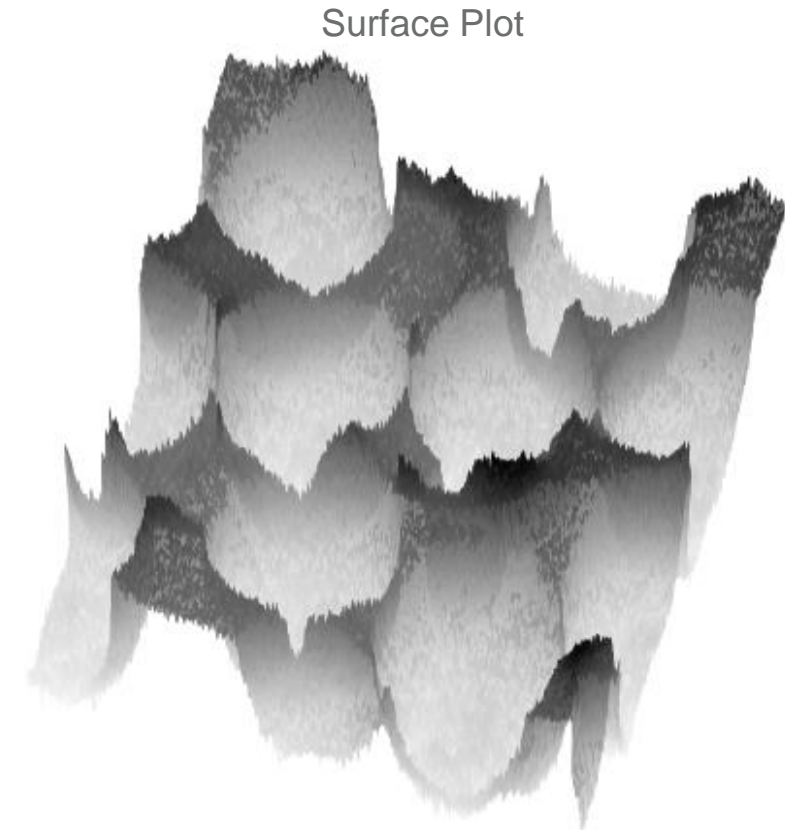
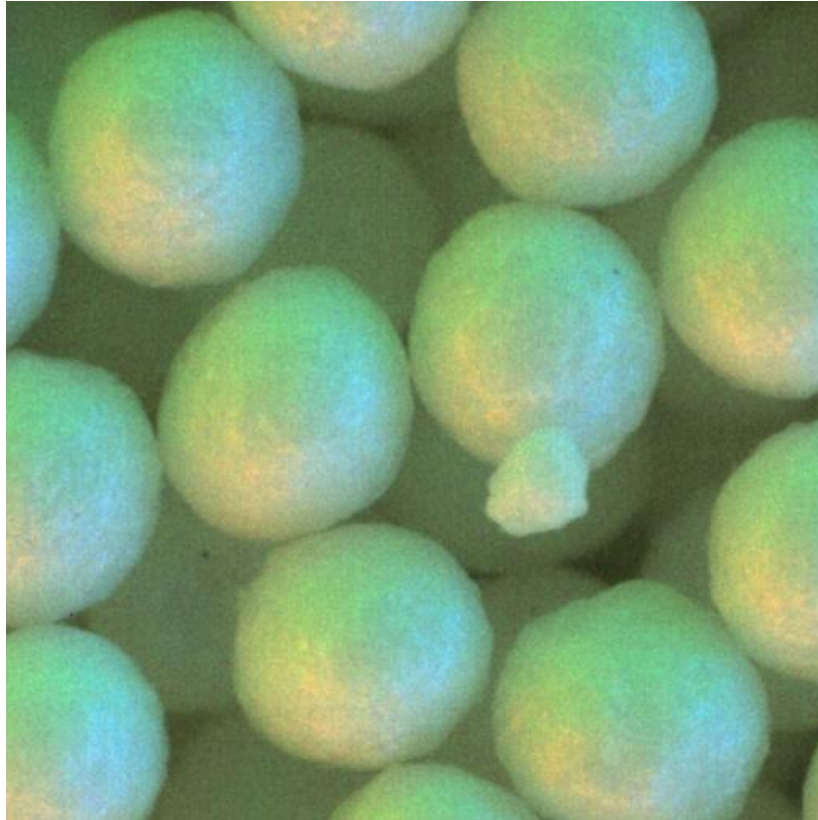
Image Segmentation



Classical Image Processing - Thresholding

-
-
-
-
-

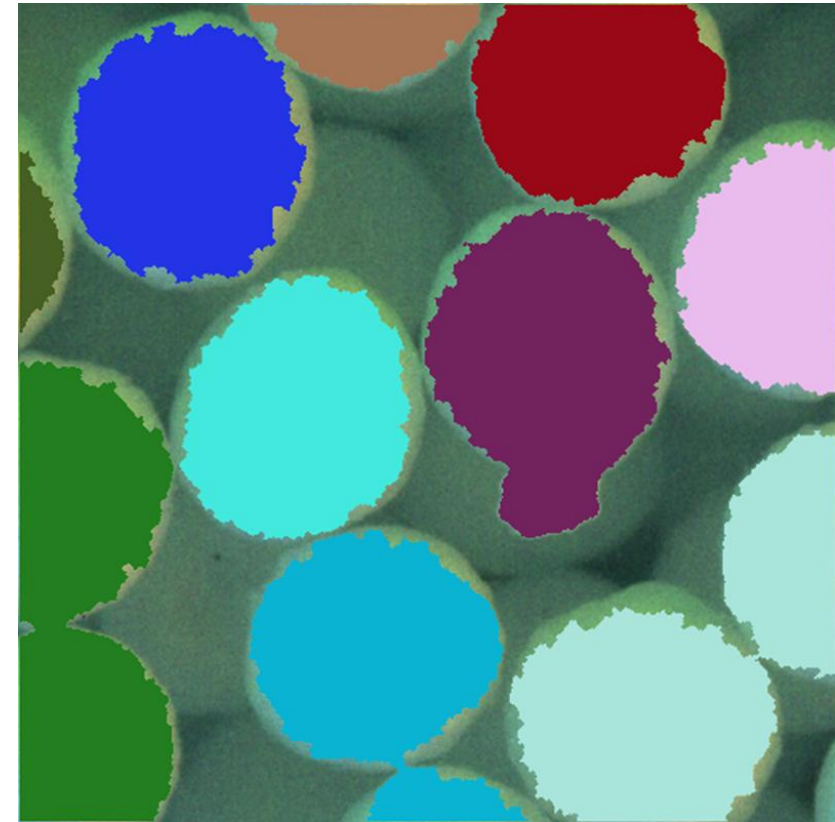
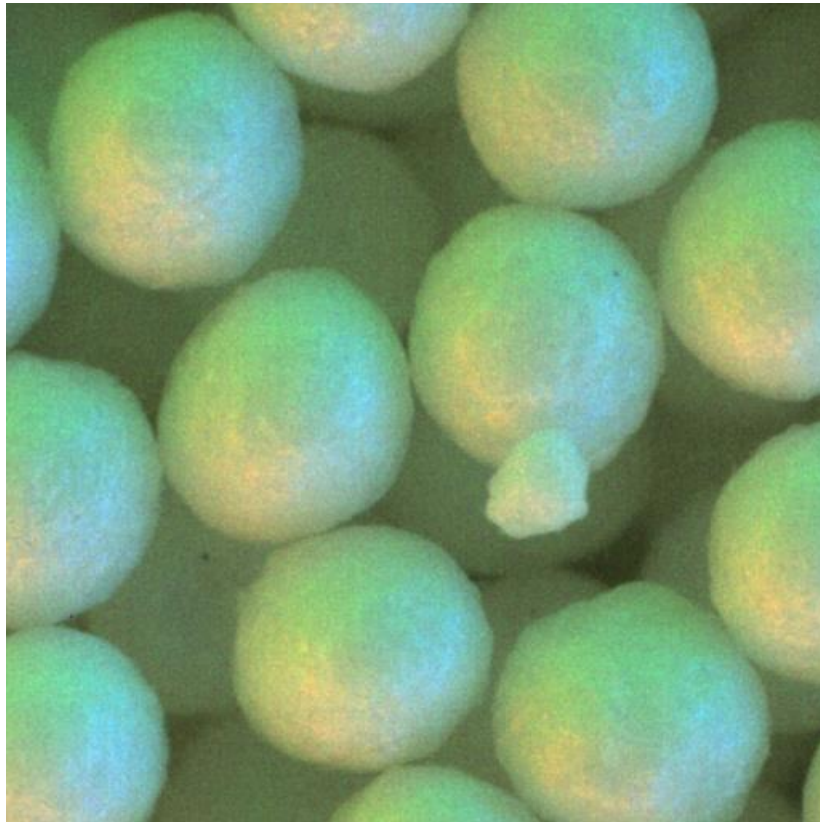
Image Segmentation



Classical Image Processing - Watershed

• • • • •

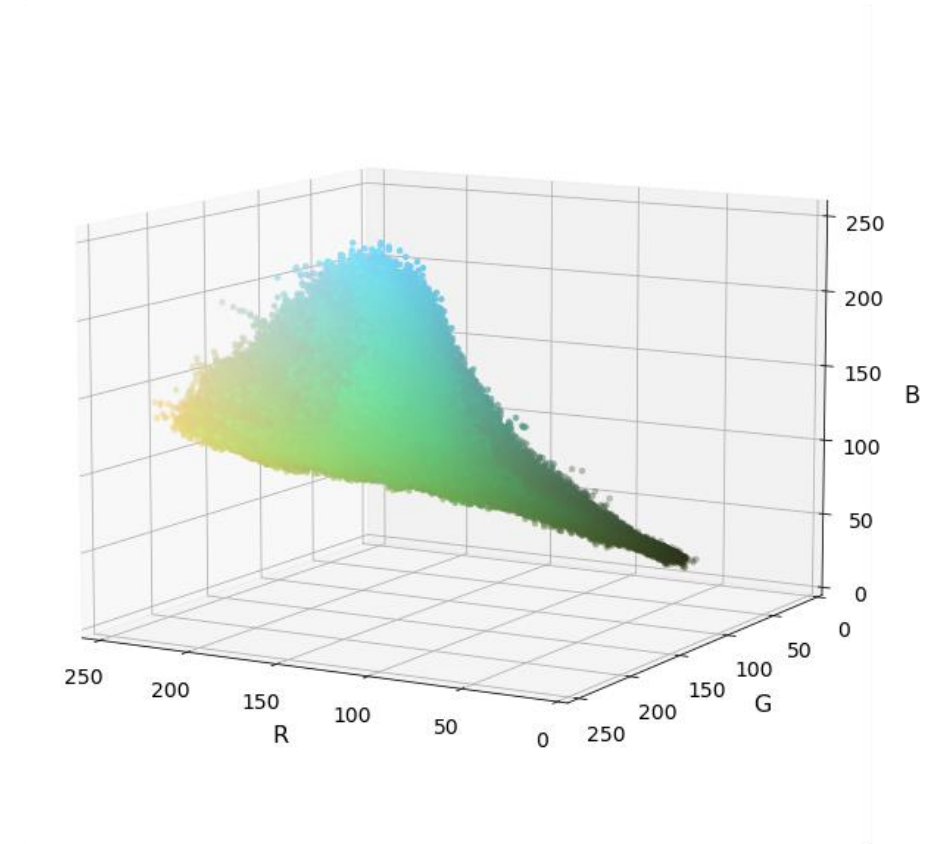
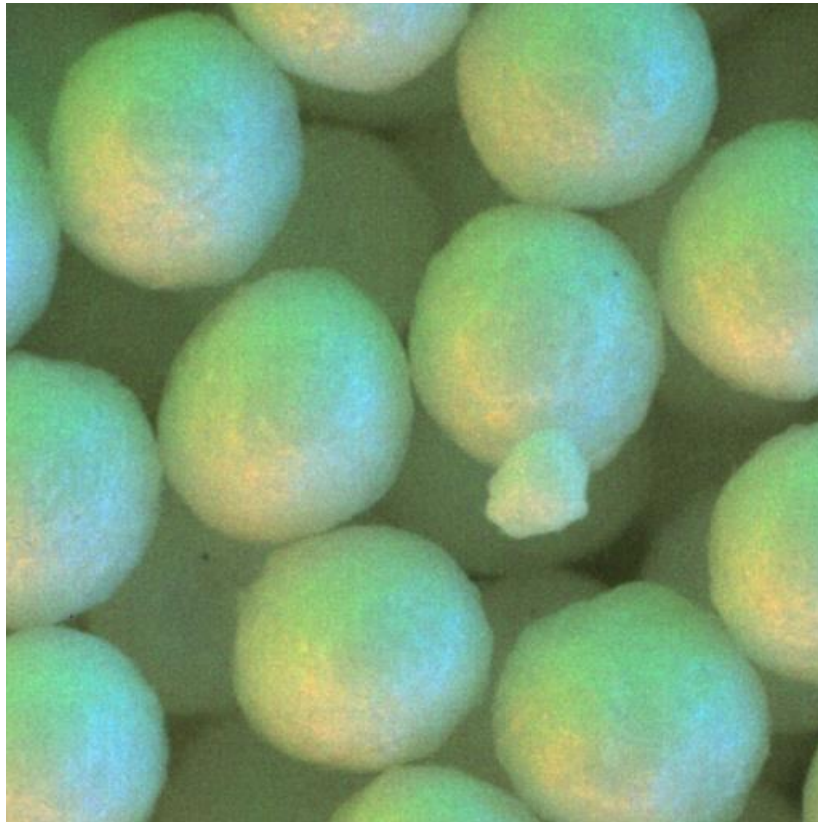
Image Segmentation



Classical Image Processing - Watershed

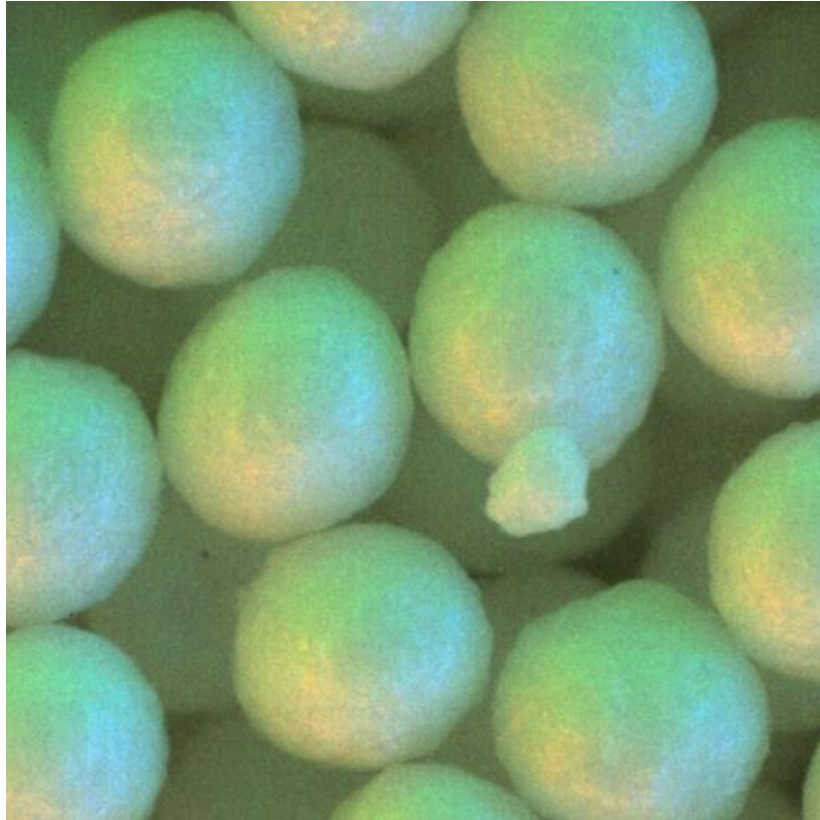
-
-
-
-
-

Image Segmentation



Machine Learning - Clustering

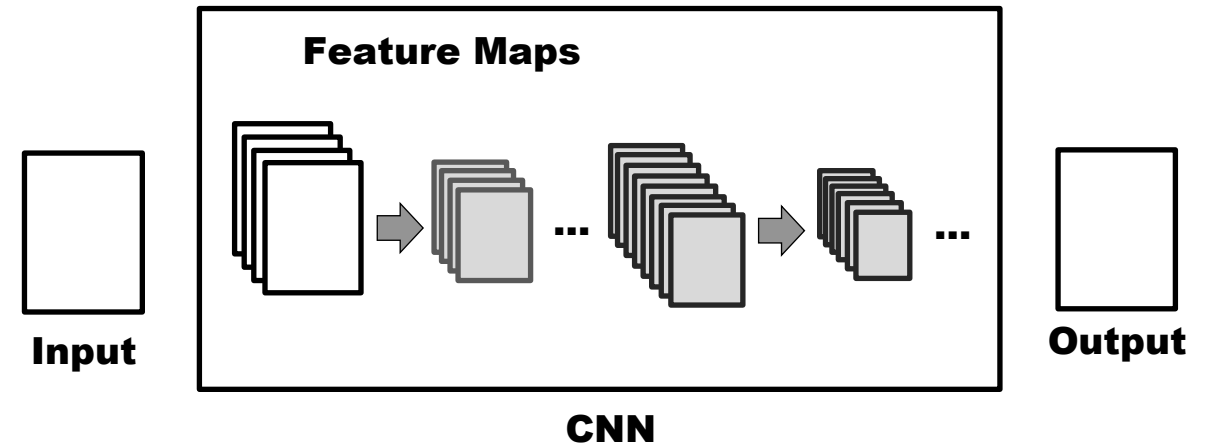
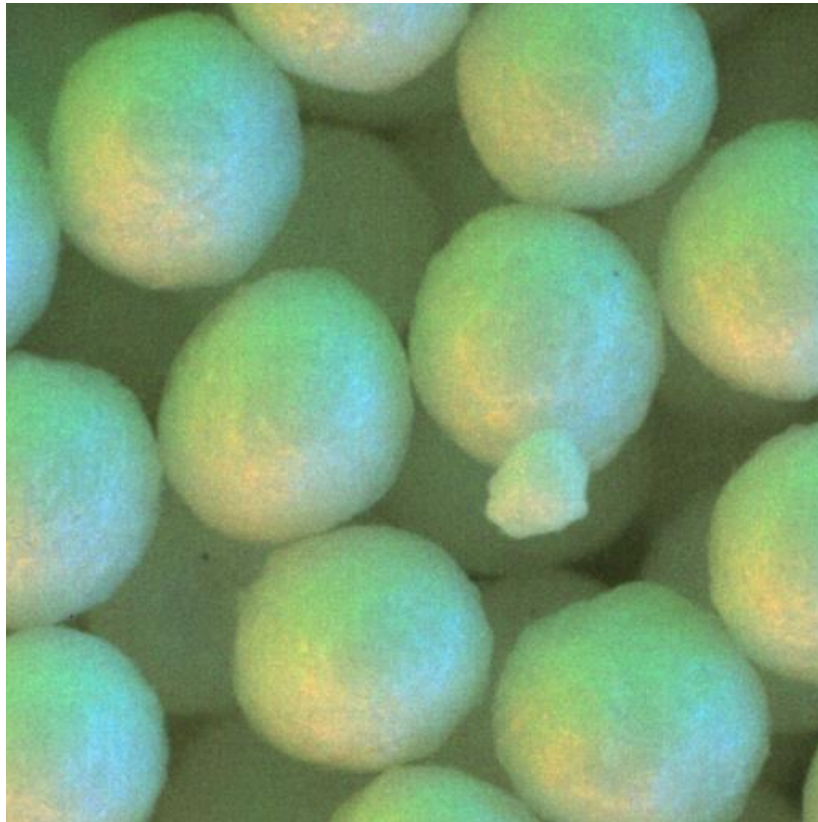
Image Segmentation



Machine Learning - Clustering

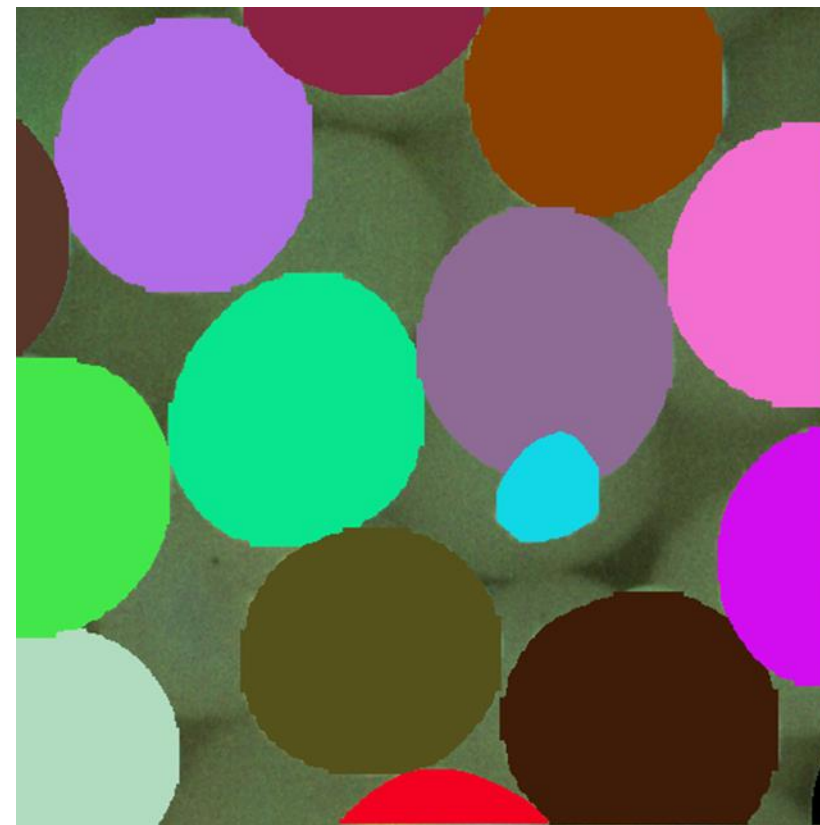
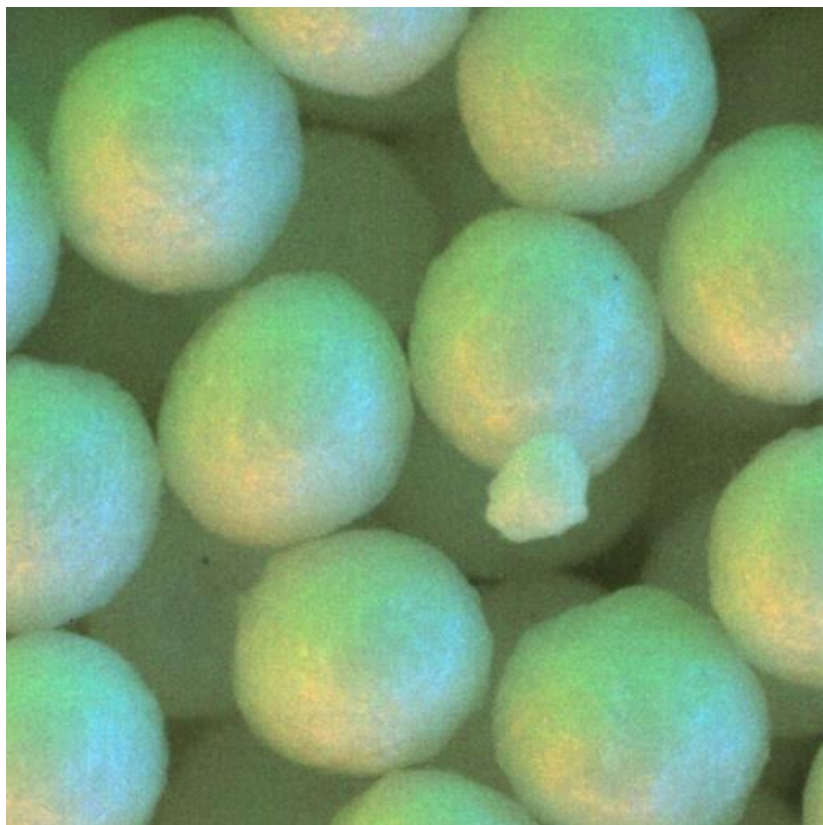
-
-
-
-
-

Image Segmentation



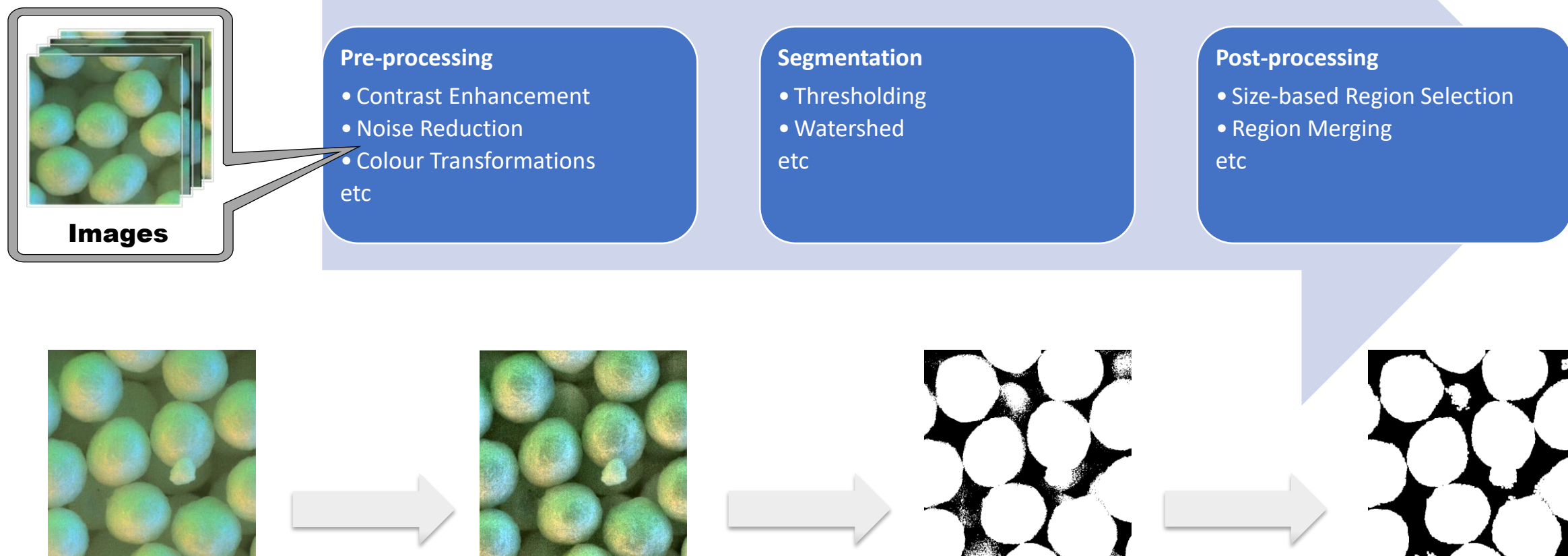
Machine Learning - Convolutional Neural Networks (CNNs)

Image Segmentation

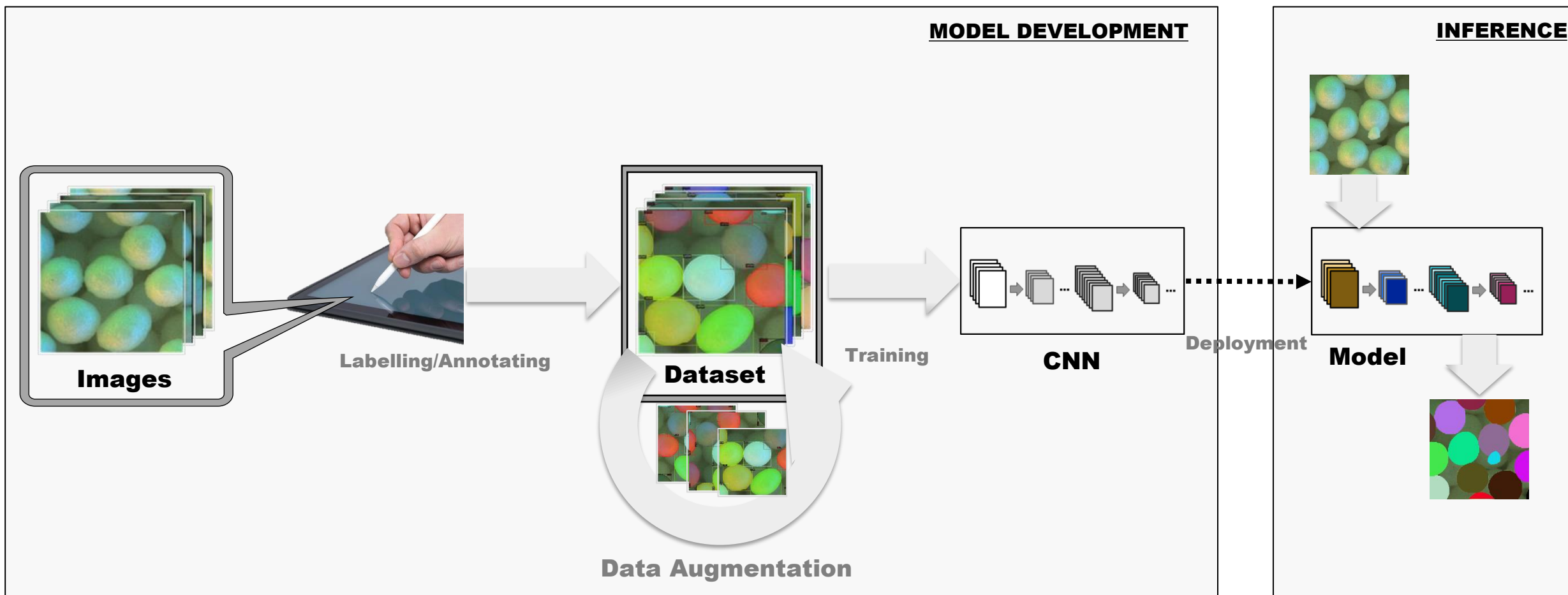


Machine Learning - Convolutional Neural Networks (CNNs)

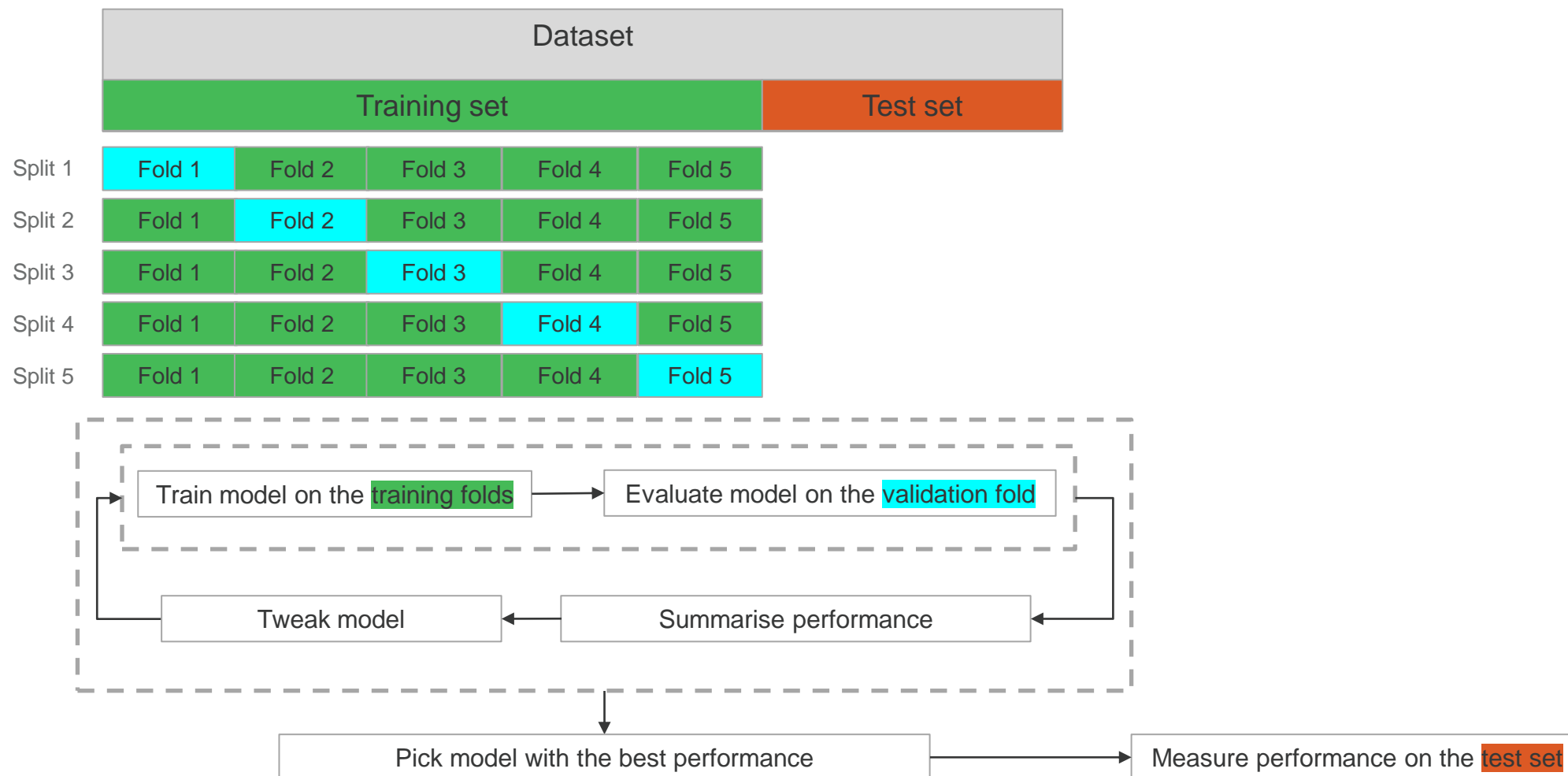
Classical Image Processing



Convolutional Neural Networks



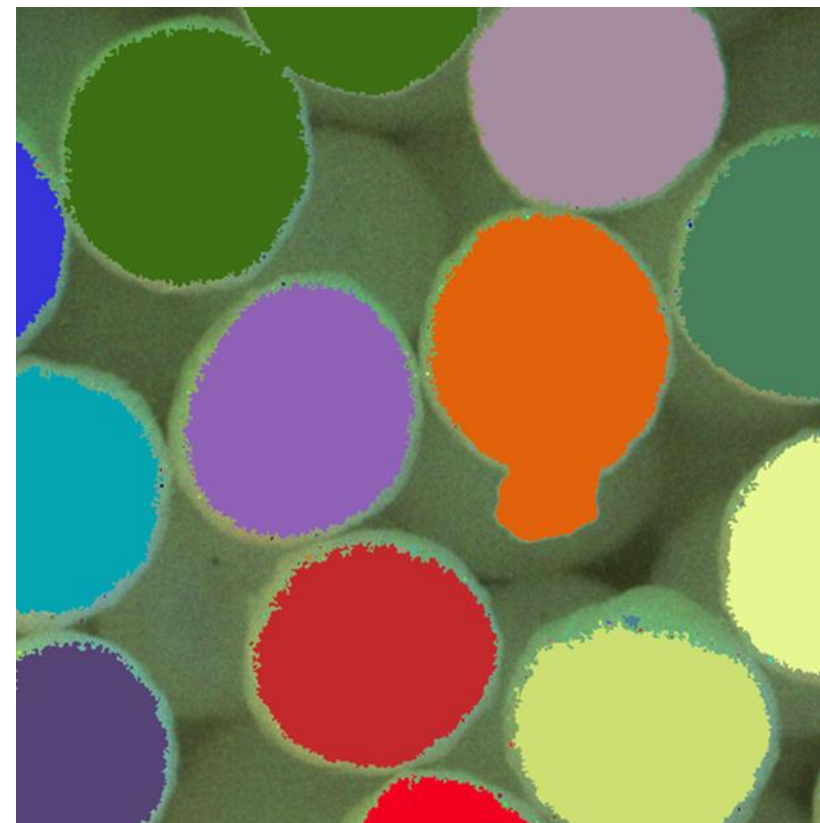
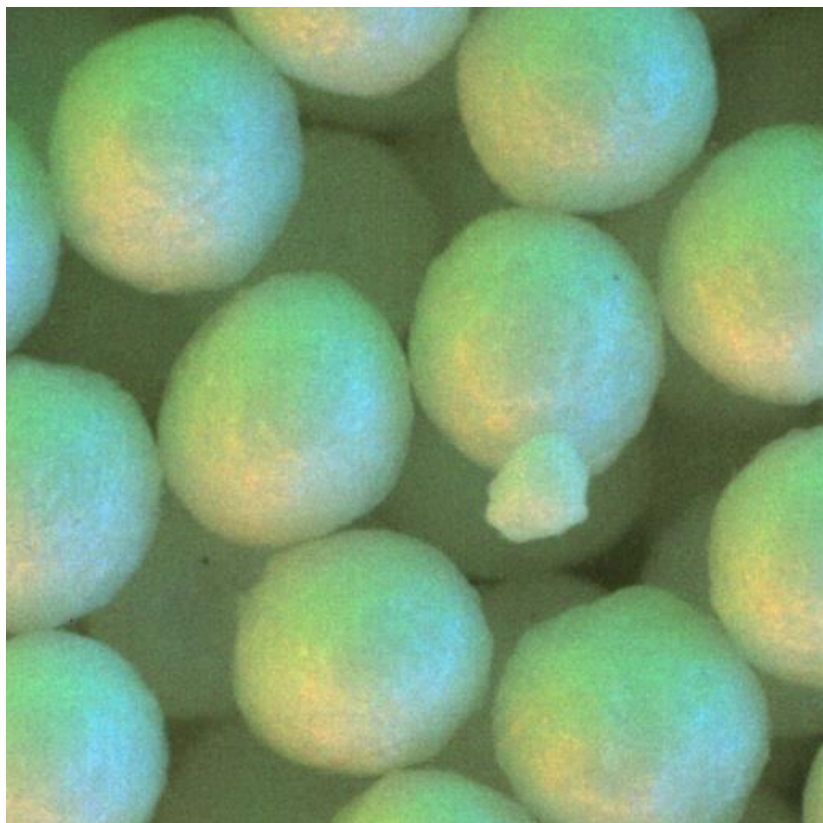
Model Development Process - K-fold cross validation



-
-
-
-
-

Image Segmentation

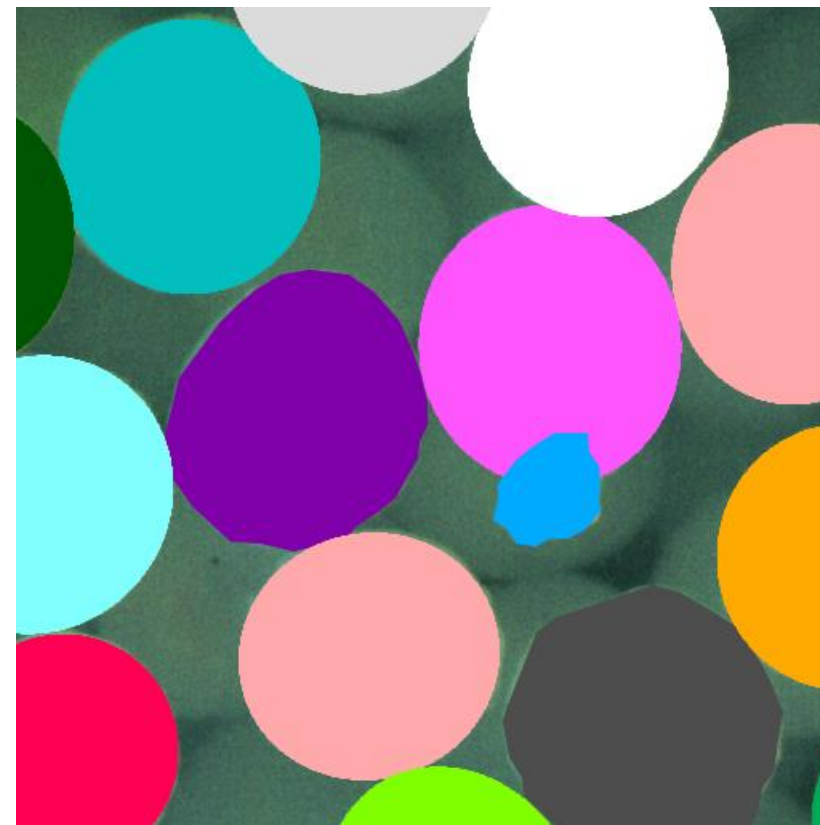
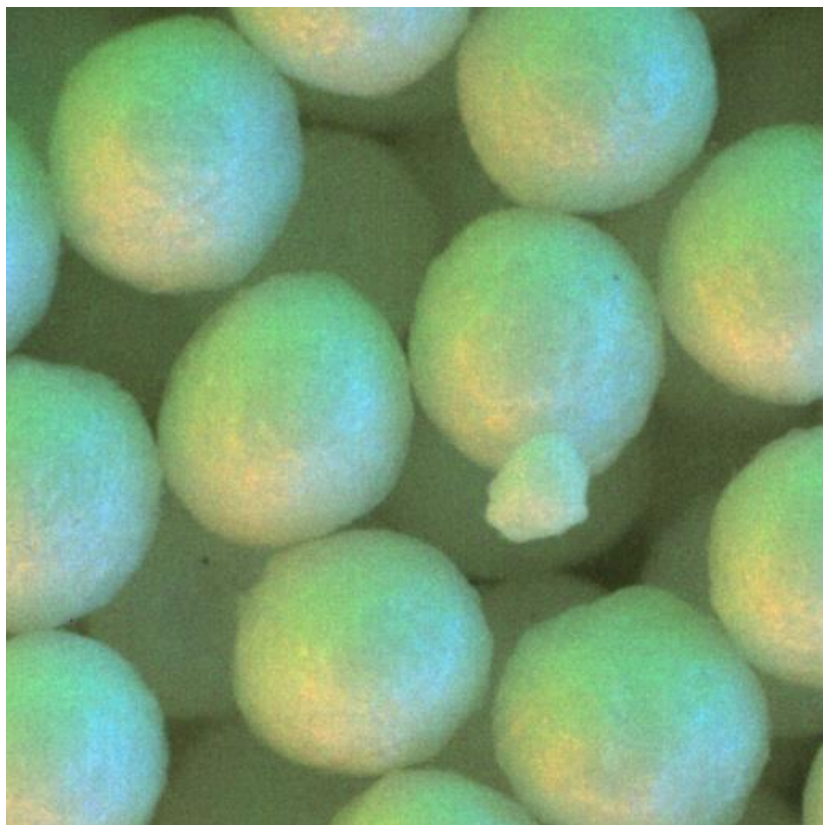
In EyePASS v2...



• • • • •

Image Segmentation

...compared to EyePASS v3



User Experience

EyePASS V2.0

Measurement Mode

In Line

Measurement Method

V2

Image Storage Interval (seconds)

15

Load Defaults for Process

Fluidised Bed Granulation

Fluidised Bed Granulation

Fluidised Bed Coating

Spheronisation

Twin Screw Granulation

Milling

Statistical Analysis

Integration Time (seconds)

60

Minimum Reported Diameter (µm)

25

Maximum Reported Diameter (µm)

1000

Image Analysis

Maximum Detection Diameter (1/4m)

1000

Minimum Detection Diameter (1/4m)

70

Average Image Intensity

0

False Agglomeration Filter

0.4

Edge Contrast Threshold

80

Analysis Block Size (pixels)

301

No. Levels

18

Exit

Save & Exit

Next >

EyePASS V3.0

Configuration: New Config

Configuration

Analysis

Camera

Alarm

Report

Measurement Mode

In Line

Measurement Method

V3

Image Storage Interval (seconds)

15

Load Defaults for Process

ML Model 001 - General

ML Model 001 - General

ML Model 002 - Glass Beads

ML Model 003 - Milling

Statistical Analysis

Integration Time (seconds)

60

Minimum Reported Diameter (µm)

25

Maximum Reported Diameter (µm)

1000

Image Analysis

Mask Threshold

0

Max Object Size px

0

Min Object Intensity

0

Max Num Objects

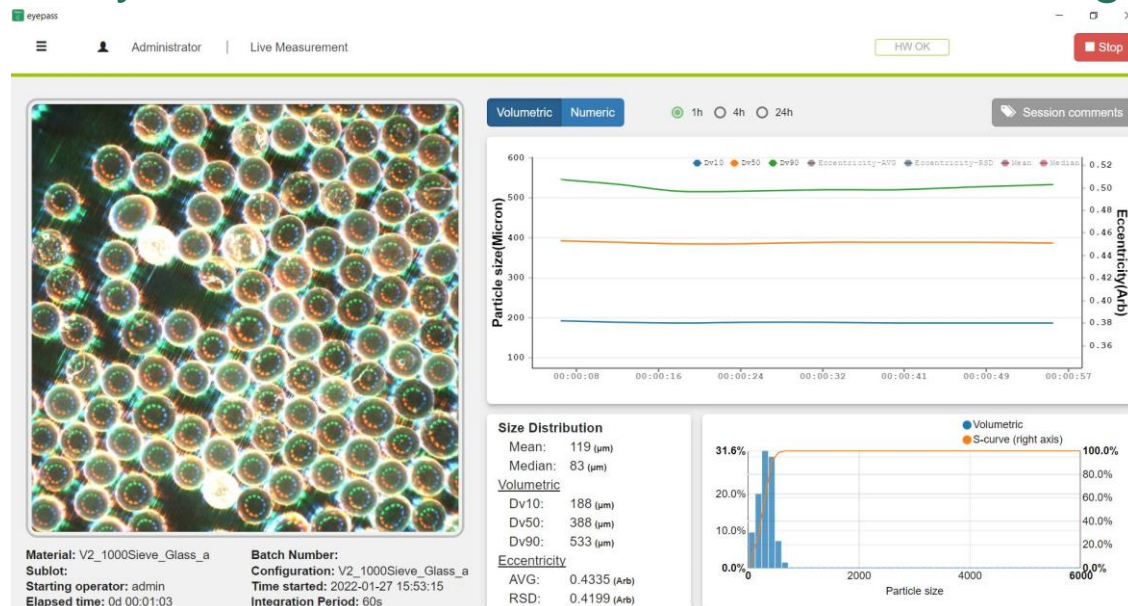
0

Exit

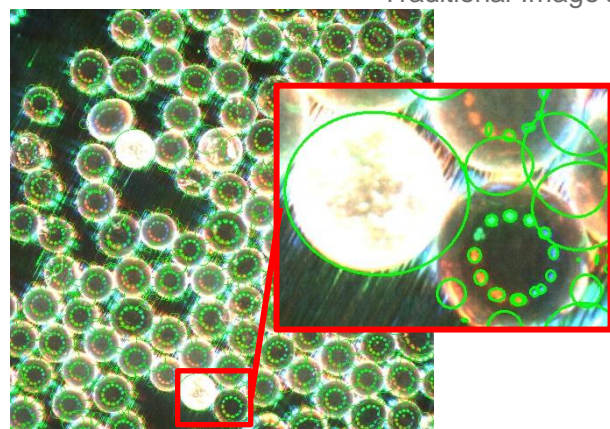
Save & Exit

Next >

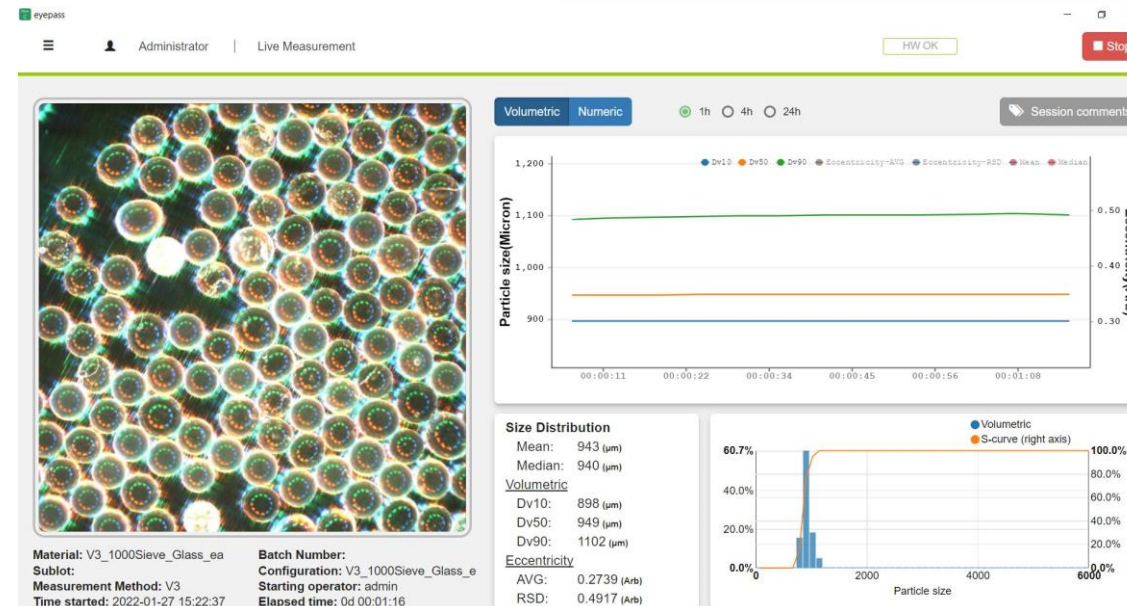
EyePASS - Before and After Machine Learning



Traditional Image analysis



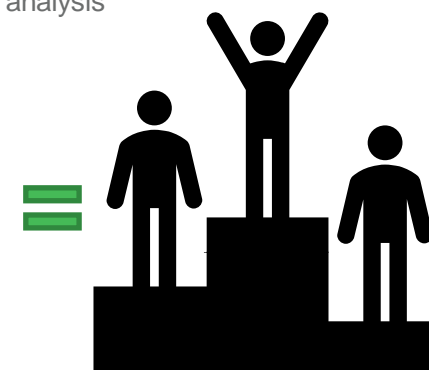
Dv10: 188 (µm)
 Dv50: 388 (µm)
 Dv90: 533 (µm)



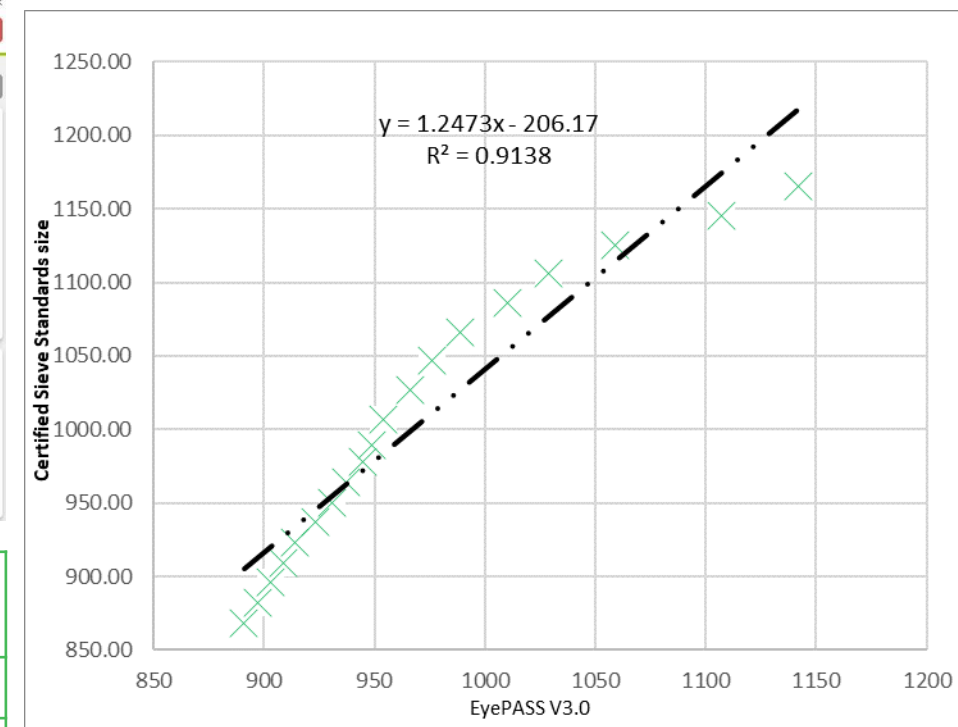
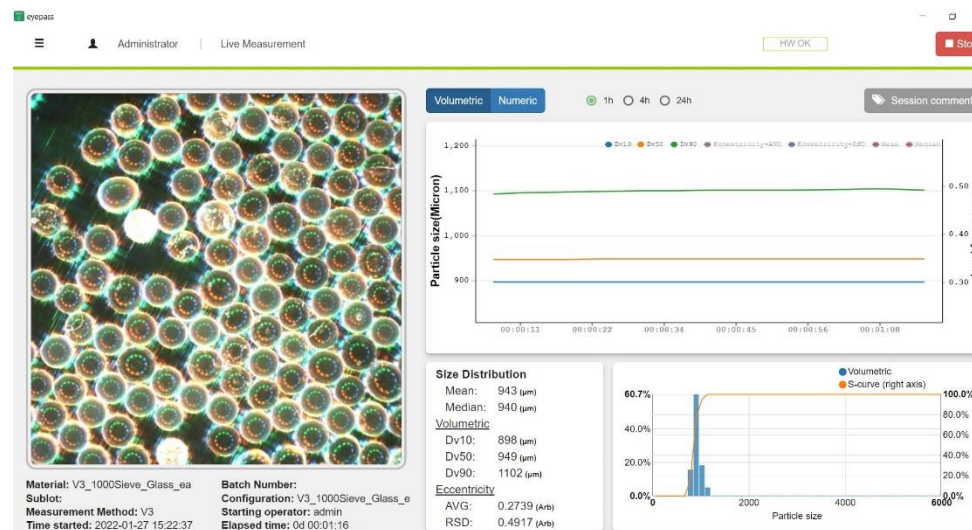
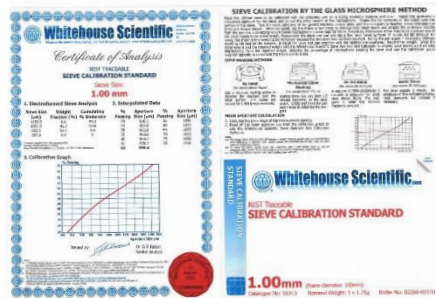
Machine Learning Image analysis



Dv10: 898 (µm)
 Dv50: 949 (µm)
 Dv90: 1102 (µm)



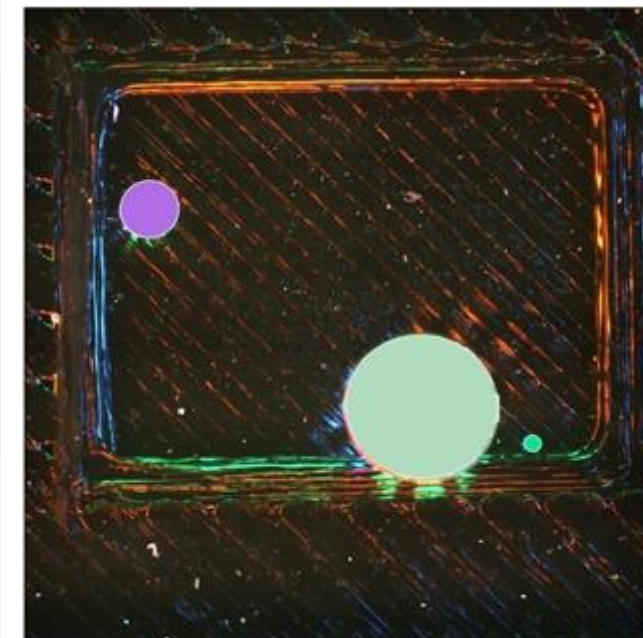
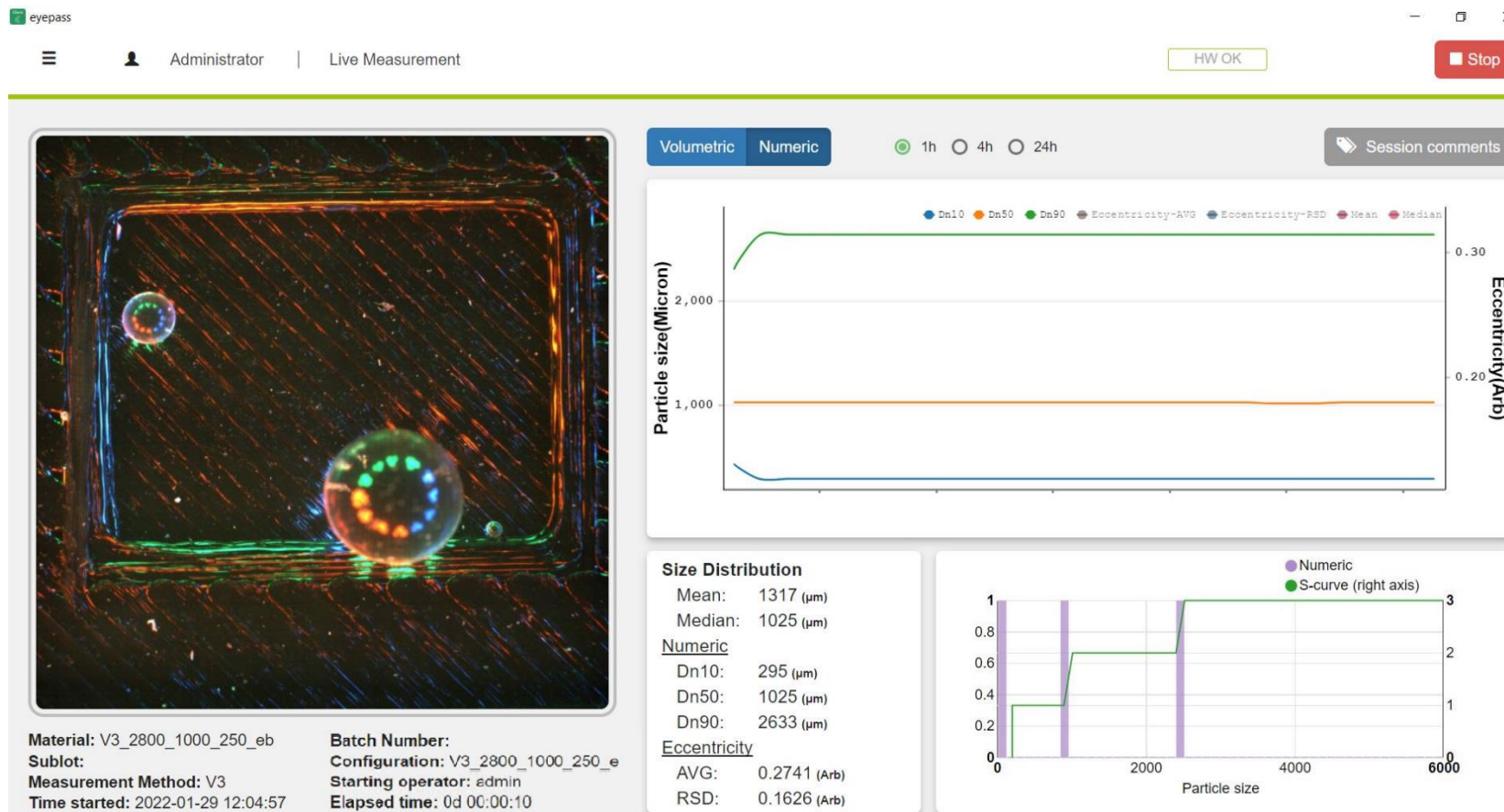
EyePASS V3.0 - Results



Correlation of 0.9559 = Strong Positive

	Glass Sieve Std um	EyePASS V3.0 (Sub sample) um	um diff	% diff
1000um D10	882.1	898	15	2 %
D50	989.6	949	41	4 %
D90	1145.6	1102	39	3 %

Untapped Potential

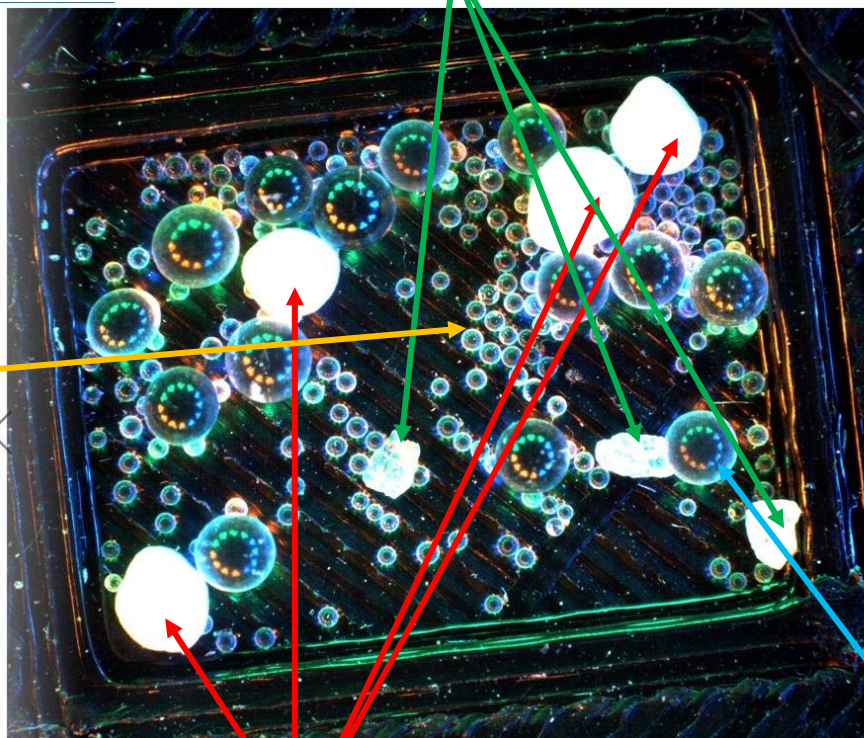


Inspiring Capability

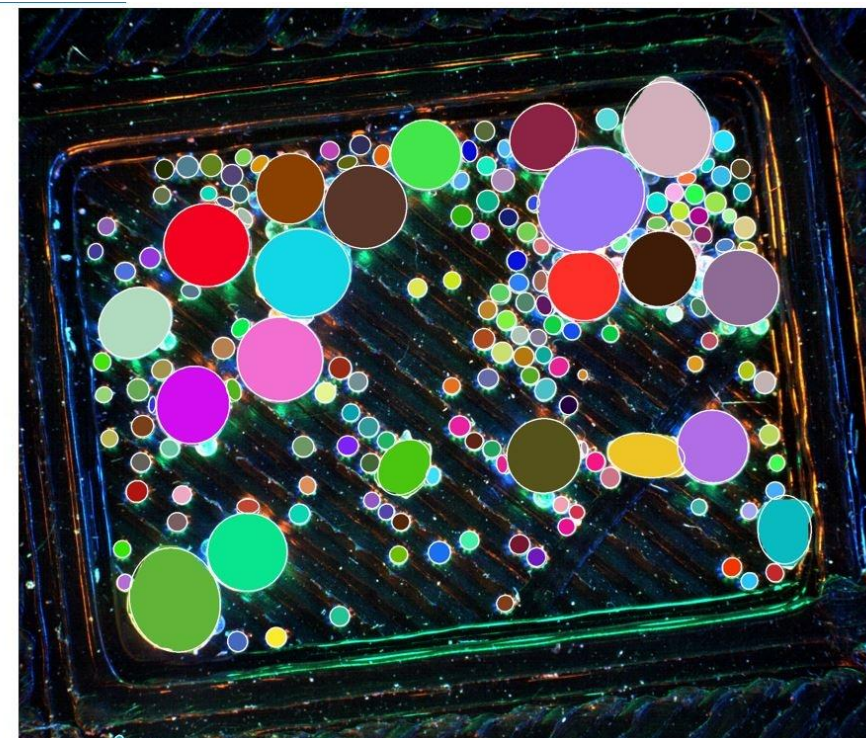
4 materials
ML
mitigates issues
like false
detections

500um Alu Oxide
(Bonded Abrasives)

250um Glass
Microspheres
(Sieve Calibration
Standard)

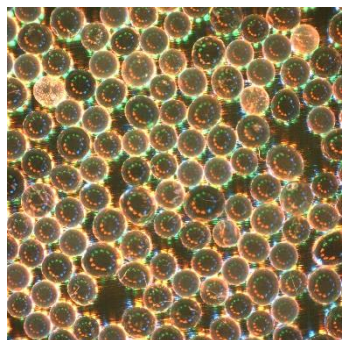


1000um Multiparticulates
(Wurster materials, cellets, suglets)

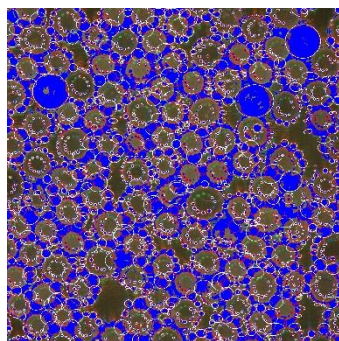


1000um Glass Microspheres
(Sieve Calibration Standard)

The Power of Machine Learning



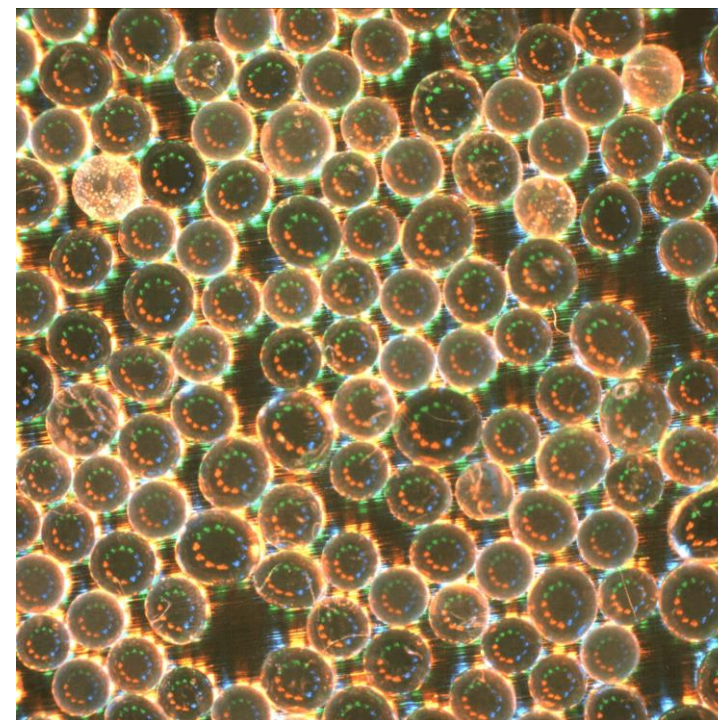
Eyecon₂ Image



EyePASS V2.0



EyePASS V3.0



Thank you!

Darren McHugh

Senior Product Manager

Innopharma Technology Ltd

mchughd@innopharmalabs.com

Girish Mallya

Senior Software Developer

Innopharma Technology Ltd

Mallyag@innopharmalabs.com