Particle size



Particle shape



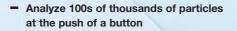
Advanced image analysis for high sensitivity particle characterization



Introducing a new concept in image analysis

The Morphologi G2 high sensitivity particle analyzer is more than just a microscope. It brings together the very best hardware and software in a single integrated package and provides the very highest level of automation and validation of results.

Morphologi



- Particle shape and count as well as size information
- Record high resolution images of every particle
- Automatic optics selection, focus and light intensity control
- Provides technical compliance with 21CFR Part 11
- Dedicated sample preparation device

The new Morphologi G2 high sensitivity particle analyzer provides repeatable and routine characterization of particle size, shape and count.

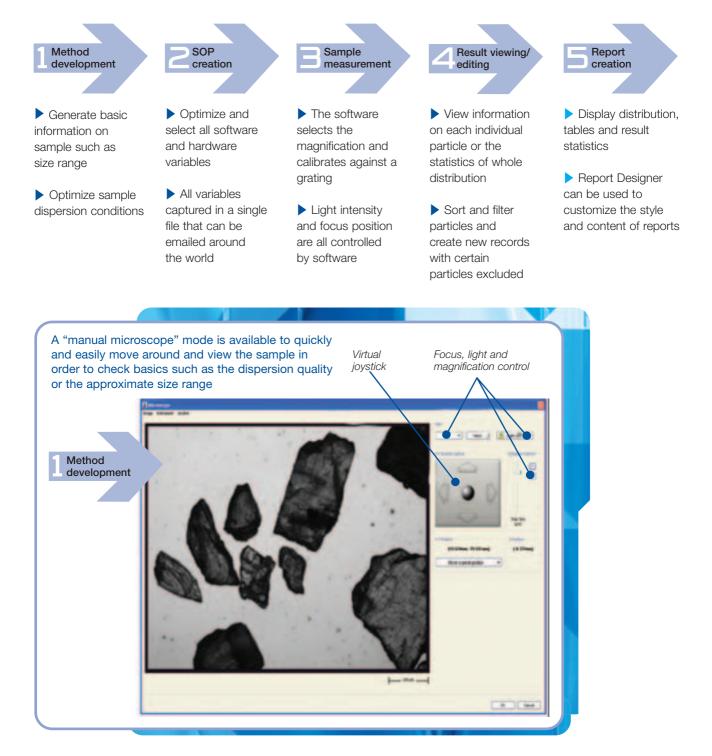
Morphologi G2 is equipped with the renowned Nikon CFI 60 optical system coupled with a high resolution digital camera for high definition aberration-free images.

Microscope-quality images and statistically significant histograms offer both qualitative and quantitative information which eliminates operator bias and saves precious man-hours.

Whether you work in R&D, process analysis or quality control, the Morphologi G2 delivers reliable, repeatable and validated results in minutes.

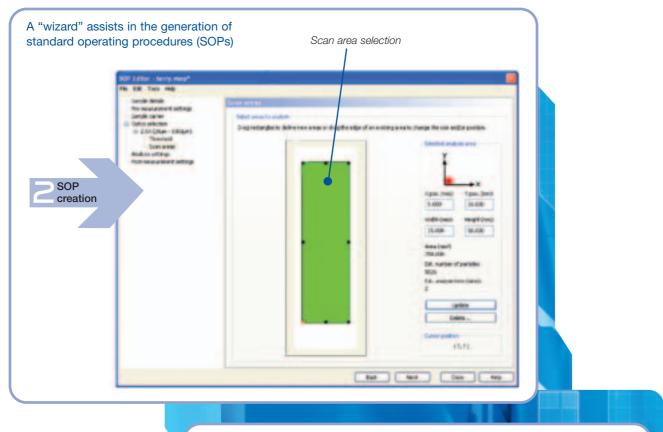


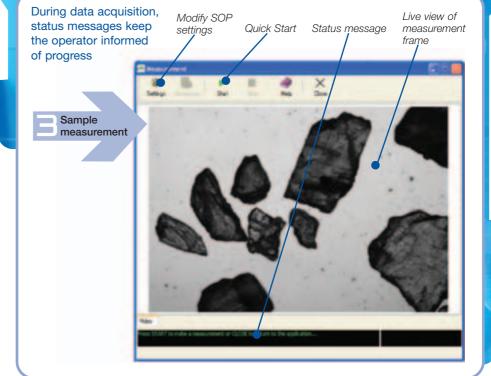
5 steps to a perfect measurement



3

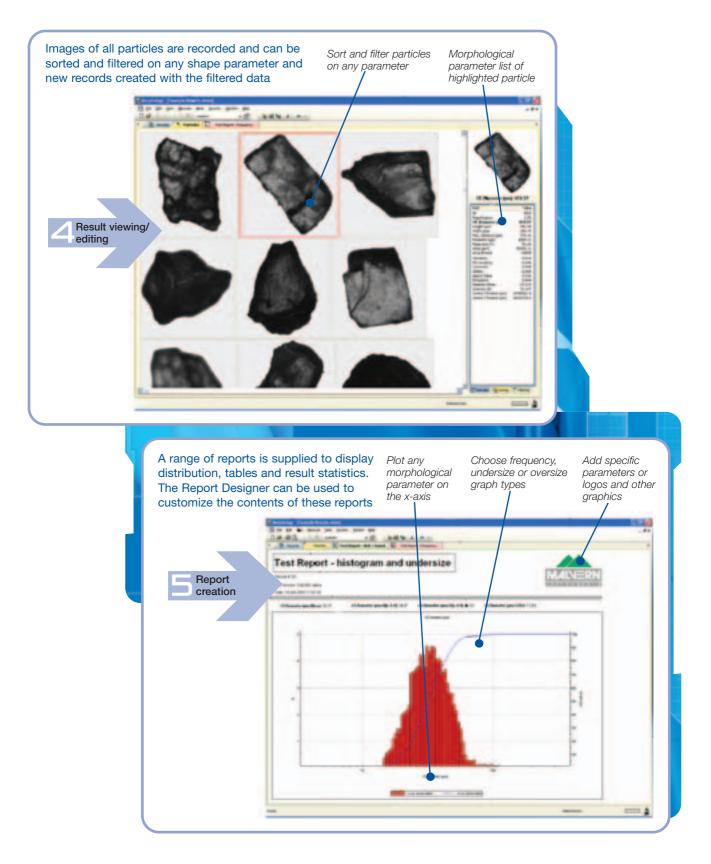






Morphologi







Length

Major atis

Why is shape analysis important?

Morphologi

Manual microscopy and traditional particle sizing techniques are often not sufficiently sensitive to distinguish subtle differences in raw materials. Batches of samples may differ by such a small amount that this difference is lost during the translation to a circle-equivalent or spherical-equivalent diameter. Calculating shape parameters like the ones shown in the list below allow even the most subtle differences to be identified and quantified.

| Parameter | Example value | Definition |
|------------------------------|---------------|--|
| ID | 516 | Unique ID of the particle - allocated in the order that the particles are detected |
| Magnification | 2.50 | Magnification used to make the measurement |
| CE diameter (µm) | 904.14 | The diameter of a circle with the same area as the particle |
| Length (µm) | 1306.35 | All possible lines from one point of the perimeter to another point on the perimeter are projected on the major axis (axis of minimum rotational energy). The maximum length of these projections is the length of the object. |
| Width (µm) | 678.54 | All possible lines from one point of the perimeter to another point on the perimeter are projected on the minor axis. The maximum length of these projections is the width of the object. |
| Max. Distance (µm) | 1318.07 | Largest distance between any two pixels in particle |
| Perimeter (µm) | 3619.42 | Actual perimeter of particle |
| Major axis° | 105.52 | Axis of minimum rotational energy |
| Area (µm²) | 371550.78 | Actual area of particle in sq. microns |
| Area (pixels) | 215018 | Number of pixels in particle |
| Circularity | 0.785 | Circumference of equivalent area circle divided by the actual perimeter of the partie = $2\sqrt{(\pi \text{ Area})/\text{Perimeter}}$ |
| HS Circularity | 0.616 | High sensitivity circularity (circularity squared) =4 π Area/perimeter ² |
| Convexity | 0.919 | Convex hull perimeter divided by actual particle perimeter |
| Solidity | 0.905 | Actual particle area divided by convex hull area |
| Aspect ratio | 0.519 | Width divided by length |
| Elongation | 0.461 | 1 - aspect ratio |
| Intensity mean | 61.310 | Average of all the greyscale values of every pixel in the particle |
| Intensity standard deviation | 29.841 | Standard deviation of all the greyscale values of every pixel in the particle |
| Centre x position (µm) | 6898271.5 | x co-ordinate of centre of mass of particle |
| Centre y position (µm) | 1797186.3 | y co-ordinate of centre of mass of particle |

Б

Width

Minor axis

Centre of mass



Why is shape analysis important?

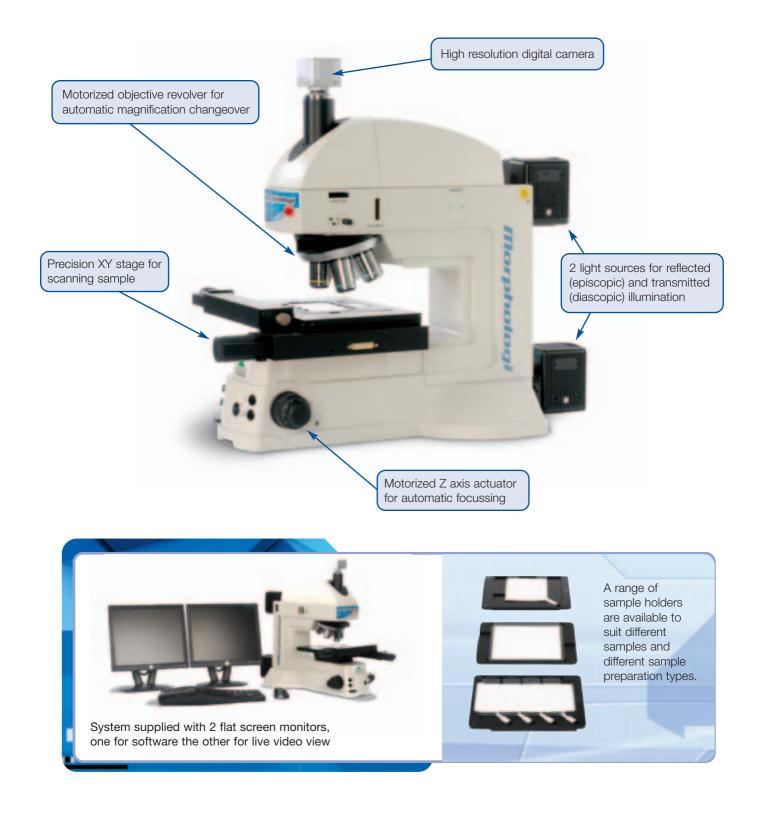
Shape parameters such as Circularity, Convexity and Elongation provide the user with a series of highly sensitive tools in order to identify and quantify subtle variations in particle shape and provide a "fingerprint" of each sample. Each parameter is usually normalized between 0 and 1 in order to provide quick and easy comparability. Traditional qualitative human descriptions such as "jagged", "smooth" or "needlelike" can be accurately quantified and hence correlated against important process or end-product variables such as flowability, active area and grinding efficiency.



| | | | | | * | |
|--|-------------|-------------|-------------|-------------|-------------|-------------|
| Circularity is a measure of the closeness to a perfect circle. | Circularity | Circularity | Circularity | Circularity | Circularity | Circularity |
| Circularity is sensitive to both changes in overall form and surface roughness. | = 1 | = 0.47 | = 0.89 | = 0.52 | = 0.47 | = 0.21 |
| Convexity is a measure of the surface roughness of a particle. Convexity is sensitive to changes in surface roughness but not overall form. | Convexity | Convexity | Convexity | Convexity | Convexity | Convexity |
| | = 1 | = 1 | = 1 | = 1 | = 0.70 | = 0.73 |
| Elongation is a measure of the length-width relationship. | Elongation | Elongation | Elongation | Elongation | Elongation | Elongation |
| Elongation is unaffected by surface roughness – a smooth ellipse has a similar elongation as a spiky ellipse of similar aspect ratio. | = 0 | = 0.82 | = 0 | = 0.79 | = 0.24 | = 0.83 |
| | | | | 1 | | 1 |



High quality hardware means high quality images



8

High quality hardware means high quality images

The Morphologi G2 is built upon the acclaimed Nikon CFI 60 optical system which achieves both higher Numerical Apertures (NA) and longer working distances than ever before. A precision XY stage and calibration grating ensure that data is precise, secure and validated at all times.

In these revolutionary optics, both axial and lateral chromatic aberration have been corrected independently in the objective and the tube lens. This geometry produces images that are crisp and clear with high contrast and minimal flare.

Morphologi

The precision engineered XY stage uses high accuracy, ground ball-screws to provide smooth and maintenance free motion with zero-backlash. The quiet and precise stepper motors ensure accurate positioning of the stage while the use of micro-stepping provides smooth motion at low speeds.

Precision etched chrome-on-glass gratings are built into the XY stage for calibration purposes. The gratings are certified and traceable to the National Physical Laboratory. The system automatically calibrates before every measurement to guarantee validated, secure data.









What Morphologi G2 delivers

| You asked for | We give you |
|---|---|
| Repeatability and automation | The tried and tested SOP (Standard Operating Procedure) methodology records all software and hardware variables in a single file. At the click of a mouse the system selects and calibrates the required magnification, the light intensity and focus before scanning a defined area. |
| Sensitivity to shape | Particles are fully characterized by morphological parameters including circle equivalent diameter, circularity and convexity. This high quality information can be used to distinguish between materials that appear identical to a conventional microscope or traditional particle sizer. |
| High quality optics | Nikon's acclaimed CFI60 optics offer longer working distances and high N.A.s and allow high contrast imaging with a minimum of flare. |
| Statistical significance | Large numbers of particles (typically 5,000 - 500,000) are captured and analyzed in minutes or even seconds. |
| Images you can see | All images are saved for future reference including the xy coordinates of each particle. If desired, you can precisely move the camera back to any position for a more detailed visual analysis. |
| Controlled orientation | To avoid errors due to random orientation, particles are dispersed onto a flat glass plate. This achieves consistency of orientation with the largest area facing the camera. |
| Regulatory compliance | The Morphologi G2 has a full validation documentation package available and provides technical compliance with the requirements of 21CFR part 11. |
| A product and company with a secure future | Recognizing that most of our customers have global operations Malvern Instruments is committed to providing a service and support structure which is present worldwide including comprehensive applications knowledge |

10



Applications and case-studies

At any point in your manufacturing process from early research and development, through process-analysis, manufacturing trouble-shooting and root-cause analysis to final product quality control, this instrument gives you an unprecedented level of product and process understanding.



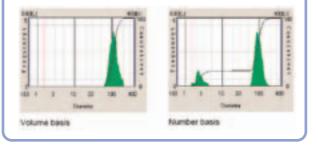
Pharmaceuticals

Even subtle differences in particle size or shape can significantly affect bioavailability, flowability,

stability, blending and tabletting efficiency. Manufacturing processing steps including crystallization, drying, milling, blending, filtering can all introduce variability into the product and have to be precisely controlled. The extra sensitivity and resolution available in the Morphologi G2 instrument provides users with the ability to identify, measure and monitor those process variables which are

Sensitivity to fines

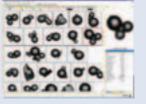
Image analysis procedes on a 'number-basis' where the contribution each particle makes to the distribution is the same - a very small particle has exactly the same weighting as a very large particle. For diagnostic or trouble-shooting purposes the presence of fines could be critical to understand any given manufacturing process.



Foreign particle detection

critical to product quality.

Image analysis is an ideal technology for detecting the presence of very small numbers of foreign particles or confirming phenomena such as agglomeration. Using single parameters or combinations of parameters, foreign particles can be detected and quantified. For example – needles or fibres can be detected using the circularity shape descriptor.





| Overview | Morphologi G2 | | | | | | |
|--|--|-------------------|--------------------|----------|-------|--|--|
| | Size, shape and count measurement of particulate samples | | | | | | |
| Size measurement | | | | | | | |
| Size range | 0.5µm - 1000µm (depending upon material properties and dispersion conditions) | | | | | | |
| Shape measurement | Multiple shape parameters calculated for each particle and distribution generated on each parameter. Parameters include: circle equivalent diameter, Length, Width, Perimeter, Area, Aspect ratio, Circularity, Convexity, Solidity, Elongation, Intensity. | | | | | | |
| Optical configurations | | | | | | | |
| Optical system | Nikon CFI 60 Brightfield/Darkfield system | | | | | | |
| Magnification (at camera) | 2.5X | 5X | 10X | 20X | 50X | | |
| Approximate total magnification (at 17" screen) | 120X | 240X | 480X | 960X | 2400X | | |
| Min particle size (µm) | 20 | 10 | 5 | 3 | 0.5 | | |
| Max particle size (µm) | 1000 | 430 | 210 | 100 | 40 | | |
| Numerical aperture | 0.075 | 0.15 | 0.30 | 0.40 | 0.55 | | |
| Focal depth (total) (µm) | 97.78 | 24.44 | 6.11 | 3.44 | 1.82 | | |
| Working distance (mm) | 8.80 | 18.00 | 15.00 | 13.00 | 9.80 | | |
| Camera system | | | | | | | |
| Camera type | 1/1/8" Global shutter progressive scan CCD | | | | | | |
| Connection protocol type | IEEE 1394a (Firewire™) | | | | | | |
| Number of pixels | 1624 x 1236 (2 MegaPixel) | | | | | | |
| Pixel size | 4.4µm x 4.4µm | | | | | | |
| Sensor size | 7.15mm x 5.44mm | | | | | | |
| Minimum PC specification (Supplied with system) | DELL Mini Tower PC, Windows XP SP2, 3.0GHz Intel Pentium IV Processor, 1Gb RAM, 160Gb-HDD, DVD +/-R/RW, complete with mouse, keyboard and 2 x 17" Flat Panel Monitors (1 for software and 1 for live video feed) | | | | | | |
| Weight and dimensions | | | | | | | |
| Weight (with stage fitted) | 50kg | | | | | | |
| Overall dimensions (with stage fitted) mm | 550(w) x 850(d) x 680(h) | | | | | | |
| Suggested deskspace (with PC and 2 screens) | 850(d) x 25 | 600(w) | | | | | |
| Site requirements | | | | | | | |
| Power requirements | AC 100-24 | 0V, 50-60Hz | | | | | |
| Ambient operating temperature | 10°C - 35°(| | | | | | |
| Humidity | 10-90% non-condensing | | | | | | |
| Location | Normal lab | oratory conditior | ns – out of direct | sunlight | | | |

Malvern Instruments Limited

Enigma Business Park • Grovewood Road • Malvern • Worcestershire • UK • WR14 1XZ **Tel:** +44 (0)1684 892456 • **Fax:** +44 (0)1684 892789

Malvern Instruments Worldwide

Sales and service centres in over 50 countries for details visit www.malvern.co.uk/contact





Malvern Instruments is part of Spectris plc, the Precision Instrumentation and Controls Company.

