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-1,25-Dihydroxyvitamin D<sub>3</sub> blocks binding and functions of nuclear vitamin D receptors of a human breast cancer cell line, MCF-7. The role of vitamin D in breast cancer development and progression is being studied intensively.

Epidemiological studies as well as laboratory-based studies on cell cultures indicate that low dietary intake of vitamin D, as well as a lack of exposure to sun light due to less time spent outdoors, may contribute to the risk of breast cancer development. Reports on the expression of vitamin D receptor (VDR) in breast cancer are not consistent. This study aimed to investigate VDR expression and the effect of

1,25-dihydroxyvitamin D<sub>3</sub> (1,25(OH)<sub>2</sub>D<sub>3</sub>) on the binding and function of nuclear VDR in the human breast cancer cell line MCF-7. It was found that MCF-7 cell express VDR. 1,25(OH)<sub>2</sub>D<sub>3</sub> decreased the binding of VDR to its specific DNA sequences.

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1,25(OH)<sub>2</sub>D<sub>3</sub> increased the expression of the osteopontin gene, as well as the expression of the osteoprotegerin gene in MCF-7 cells. MCF-7 cells were also capable of secreting osteopontin, an effect inhibited by 1,25(OH)<sub>2</sub>D<sub>3</sub>. 1,25(OH)<sub>2</sub>D<sub>3</sub> inhibited the expression of matrix metalloproteinase 2 (MMP-2) by MCF-7 cells. The results obtained show that the effects of 1,25(OH)<sub>2</sub>D<sub>3</sub> on MCF-7 cells are similar to previously described effects of 1,25(OH)<sub>2</sub>D<sub>3</sub> on osteoblasts and suggest that the cancer-osteoporosis link may be due to interference with the gene expression of proteins involved in bone remodeling.

Chris Wilder's long-awaited appointment as Head Coach of Villa is not expected to be made until next week. The Sky Blues have yet to make an appointment after the Coventry side opted to part company

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AdobeRootCA.pem AdobeRootCA2.pem  
AdobeRootCA3.pem This invention relates  
generally to speed shift control of vehicle engines,  
and more particularly to a speed shift control  
system that is provided with multiple control  
modes. In vehicles powered by gasoline engines,  
engine speed is governed for different driving  
conditions to avoid engine stall or surge caused  
by an over-speed of the engine. Engines powered  
by a drive axle, such as those used for heavy-duty  
vehicles, usually include an over-speed protection  
device that reduces the engine speed when the  
engine speed is above a predetermined over-  
speed level to prevent engine stall or surge. In  
heavy-duty vehicles, an over-speed protection  
device is usually provided only when the drive  
axle is driven in vehicles equipped with the  
device. If the drive axle is not driven, or if a drive  
axle of a vehicle equipped with such a device  
malfunctions, the over-speed protection device  
does not function. That is, when the drive axle is  
not driven, the engine speed can be controlled  
only by a transmission controlled over-speed  
protection device. This makes the engine speed  
difficult to control because the engine speed can

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exceed the predetermined over-speed level to stall or surge, without any restriction. Japanese Patent Laid-Open Publication No. 2000-54240 discloses a speed shift control device that operates based on external inputs, such as road speed and vehicle speed. This publication does not include any direct reference to engine speed. Japanese Patent Laid-Open Publication No. 2001-237143 discloses a control device for controlling a shift control mechanism in a vehicle drive train. The shift control mechanism is switched to a first control mode when the vehicle speed is higher than a predetermined speed. The control device controls shift control based on speed reference data, such as road speed and vehicle speed. This control device also fails to provide any direct reference to engine speed. As the US continues to issue

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